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<p>(21) International Application Number: <b>PCT/US99/20104</b></p> <p>(22) International Filing Date: <b>3 September 1999 (03.09.99)</b></p> <p>(30) Priority Data:              60/106,344                      30 October 1998 (30.10.98)      US              60/122,541                      2 March 1999 (02.03.99)              US</p> <p>(71) Applicant (for all designated States except US): <b>WALTER REED ARMY INSTITUTE OF RESEARCH [US/US]; 503 Robert Grant Road, Bldg. 503, Room 1W38, Silver Spring, MD 20910 (US).</b></p> <p>(72) Inventors; and</p> <p>(75) Inventors/Applicants (for US only): <b>BALKIN, Thomas, J. [US/US]; 3839 Parrot Drive, Ellicott City, MD (US). BELENKY, Gregory, L. [US/US]; 9829 La Duke Drive, Kensington, MD 20895 (US). HALL, Stanley, W. [US/US]; 15352 Baileys Lane, Silver Spring, MD 20906 (US). KAMIMORI, Gary, H. [US/US]; 14918 Cherrywood Drive, Laurel, MD 20707 (US). REDMOND, Daniel, P. [US/US]; 10704 Bucknell Drive, Silver Spring, MD 20902 (US). SING, Helen, C. [US/US]; 905 Sligo Creek Parkway, Takoma Park, MD 20912 (US). THOMAS, Maria, L. [US/US]; 9522 Caboose Court, Columbia, MD 21045-3208 (US). THORNE, David, R. [US/US]; 1855 Calvert Street</b></p>		<p>N.W., Washington, DC 20009-1966 (US). WESENSTEN, Nancy, Jo [US/US]; 10212 Procter Street, Silver Spring, MD 20901 (US).</p> <p>(74) Agent: <b>HARRIS, Charles, H.; Cahn &amp; Samuels, LLP, Suite 200, 2000 P Street N.W., Washington, DC 20036 (US).</b></p> <p>(81) Designated States: <b>AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).</b></p> <p style="text-align: center;">Published With international search report.</p>
<p>(54) Title: <b>SYSTEM AND METHOD FOR PREDICTING HUMAN COGNITIVE PERFORMANCE USING DATA FROM AN ACTIGRAPH</b></p>		
<p>(57) Abstract</p> <p>A system and a method for providing a determination of predicted cognitive performance of an individual based the time of day and on factors including sleep history based on activity data form an actigraph. The system and method provide a numerical representation of the predicted cognitive performance. Both may be used to optimize the work schedule of the actigraph wearer to maximize the cognitive capacity during working hours.</p>		

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settings, both civilian and military, sleep deprivation reduces productivity (output of useful work per unit of time) on cognitive tasks.

Thus, using computer-based cognitive performance tests, it has been shown that total sleep deprivation degrades human cognitive performance by approximately 25% for each successive period of 24 hours awake. However, it also has been shown that even small amounts of sleep reduce the rate of sleep loss-induced cognitive performance degradation: Belenky et al. in their article entitled "Sustaining Performance During Continuous Operations: The U.S. Army's Sleep Management System," published in *20<sup>th</sup> Army Science Conference Proceedings*, vol. 2, pp. 657-661 (1996), disclose that a single 30-minute nap every 24 hours reduces the rate of cognitive performance degradation to 17% per day over 85 hours of sleep deprivation. This suggests that recuperation of cognitive performance during sleep accrues most rapidly early in the sleep period. No other factor besides the amount of sleep contributes so substantially and consistently to the normal, daily variations in cognitive performance.

In addition to sleep/wake history, an individual's cognitive performance at a given point in time is determined by the time of day. In the early 1950s, Franz Halberg and associates observed a 24-hour periodicity in a host of human physiologic (including body temperature and activity), hematologic, and hormonal functions, and coined the term 'circadian' (Latin for 'about a day') to describe this cyclic rhythm. Halberg showed that most noise in experimental data came from comparisons of data sampled at different times of day.

When humans follow a nocturnal sleep/diurnal wake schedule (for example, an 8-hour sleep/16-hour wake cycle, with nightly sleep commencing at approximately midnight), body temperature reaches a minimum (trough) usually between 2:00 AM and 6:00 AM. Body temperature then begins rising to a maximum (peak) usually between 8:00 PM and 10:00 PM. Likewise, systematic studies of daily human cognitive performance rhythms show that speed of responding slowly improves across the day to reach a maximum in the evening (usually between 8:00 PM and 10:00 PM) then dropping more rapidly to a minimum occurring in the early morning hours (usually between 2:00 AM and 6:00 AM). Similar but somewhat less consistent rhythms have been shown from testing based on various cognitive performance tasks. Thus, superimposed on the effect of total

performance decline (i.e., sleepiness as measured by MSLT increases). However, although there is a correlation between MSLT-determined sleepiness/alertness and cognitive performance (greater sleepiness as indexed by MSLT corresponding to poorer cognitive performance), this correlation has never been shown to be perfect and for the most part is not strong. As a result, the MSLT is a poor (i.e., unreliable) predictor of cognitive performance.

Subjective measures of sleepiness/alertness also have been shown to correlate (albeit weakly) with cognitive performance. Hoddes et al., in their article entitled "Quantification of Sleepiness: A New Approach" published in *Psychophysiology*, vol. 10, pp. 431-436 (1973) describe the Stanford Sleepiness Scale (SSS), a subjective questionnaire used widely to measure sleepiness/alertness. In the SSS, individuals rate their current level of sleepiness/alertness on a scale from 1 to 7, with 1 corresponding to the statement, "feeling active and vital; alert; wide awake" and 7 corresponding to the statement "almost in reverie; sleep onset soon; losing struggle to remain awake." Higher SSS scores indicate greater sleepiness. As with the MSLT, during both total and partial sleep deprivation, scores on the SSS increase. However, as with MSLT, the correspondence between SSS-determined sleepiness/alertness and cognitive performance decrements is weak and inconsistent. As a result, the SSS also is a poor predictor of cognitive performance. Some other examples of subjective measures of sleepiness/alertness include the Epworth Sleepiness Scale described by Johns in his article entitled "Daytime Sleepiness, Snoring, and Obstructive Sleep Apnea" published in *Chest*, vol. 103, pp. 30-36 (1993), and the Karolinska Sleepiness scale described by Akerstedt and Gillberg in their article entitled "Subjective and Objective Sleepiness in the Active Individual" published in *International Journal of Neuroscience*, vol. 52, pp. 29-37 (1990). The correspondence between these subjective measures and cognitive performance also is weak and inconsistent.

In addition, factors modifying cognitive performance may not correspondingly affect objective or subjective measures of sleepiness/alertness, and vice versa. For example, the Penetar et al. article entitled "Amphetamine Effects on Recovery Sleep Following Total Sleep Deprivation" published in *Human Psychopharmacology*, vol. 6, pp. 319-323 (1991), disclose that during sleep deprivation, the stimulant drug d-amphetamine improved cognitive performance but

not sleepiness/alertness (as measured by the MSLT). In a similar study, caffeine given as a sleep deprivation countermeasure maintained elevated cognitive performance for over 12 hours while the effects on subjective sleepiness, vigor and fatigue transiently improved but then decayed. Thorne et al. in their article entitled

5 "Plumbing Human Performance Limits During 72 hours of High Task Load" in Proceedings of the 24<sup>th</sup> DRG Seminar on the Human as a Limiting Element in Military Systems, Defense and Civil Institute of Environmental Medicine, pp. 17-40 (1983), describe how cognitive performance continues to decline over 72 hours of sleep deprivation whereas subjective sleepiness/alertness declined over the first 24

10 hours but subsequently leveled off. The findings that cognitive performance and measures of sleepiness/alertness are not always affected in the same way indicate that they are not interchangeable. That is, measures of sleepiness/alertness cannot be used to predict cognitive performance, and vice versa.

Methods and apparatuses related to alertness detection fall into five basic

15 categories: a method/apparatus for unobtrusively monitoring current alertness level; a method/apparatus for unobtrusively monitoring current alertness level and providing a warning/alarm to the user of decreased alertness and/or to increase user's alertness level; a method/apparatus for monitoring current alertness level based on the user's responses to some secondary task possibly with an alarm

20 device to warn the user of decreased alertness and/or to increase user's alertness level; methods to increase alertness; and a method/apparatus for predicting past, current, or future alertness.

These methods and apparatuses that unobtrusively monitor the current alertness level are based on an "embedded measures" approach. That is, such

25 methods infer alertness/drowsiness from the current level of some factor (e.g., eye position or closure) assumed to correlate with alertness/drowsiness. Some recently issued patents of this type include U.S. Patent No. 5,689,241 to J. Clarke, Sr., et al. disclosing an apparatus to detect eye closure and ambient temperature around the nose and mouth; U.S. Patent No. 5,682,144 to K. Mannik disclosing an

30 apparatus to detect eye closure; and U.S. Patent No. 5,570,698 to C. Liang et al. disclosing an apparatus to monitor eye localization and motion to detect sleepiness. An obvious disadvantage of these types of methods and apparatuses is that the measures are likely detecting sleep onset itself rather than small decreases in alertness.

In some patents, methods for embedded monitoring of alertness/drowsiness are combined with additional methods for signaling the user of decreased alertness and/or increasing alertness. Recently issued patents of this type include U.S. Patent No. 5,691,693 to P. Kithil describing a device that senses a vehicle operator's head position and motion to compare current data to profiles of "normal" head motion and "impaired" head motion. Warning devices are activated when head motion deviates from the "normal" in some predetermined way. U.S. Patent No. 5,585,785 to R. Gwin et al. describes an apparatus and a method for measuring total handgrip pressure on a steering wheel such that an alarm is sounded when the grip pressure falls below a predetermined "lower limit" indicating drowsiness. U.S. Patent No. 5,568,127 to H. Bang describes a device for detecting drowsiness as indicated by the user's chin contacting an alarm device, which then produces a tactile and auditory warning. U.S. Patent No. 5,566,067 to J. Hobson et al. describes a method and an apparatus to detect eyelid movements. A change in detected eyelid movements from a predetermined threshold causes an output signal/alarm (preferably auditory). As with the first category of methods and apparatuses, a disadvantage here is that the measures are likely detecting sleep onset itself rather than small decreases in alertness.

Other alertness/drowsiness monitoring devices have been developed based on a "primary/secondary task" approach. For example, U.S. Patent No. 5,595,488 to E. Gozlan et al. describes an apparatus and a method for presenting auditory, visual, or tactile stimuli to an individual to which the individual must respond (secondary task) while performing the primary task of interest (e.g., driving). Responses on the secondary task are compared to baseline "alert" levels for responding. U.S. Patent No. 5,259,390 to A. MacLean describes a device in which the user responds to a relatively innocuous vibrating stimulus. The speed to respond to the stimulus is used as a measure of the alertness level. A disadvantage here is that the apparatus requires responses to a secondary task to infer alertness, thereby altering and possibly interfering with the primary task.

Other methods exist solely for increasing alertness, depending upon the user to self-evaluate alertness level and activate the device when the user feels drowsy. An example of the latter is U.S. Patent No. 5,647,633 and related patents to M. Fukuoka in which a method/apparatus is described for causing the user's seat to vibrate when the user detects drowsiness. Obvious disadvantages of such

devices are that the user must be able to accurately self-assess his/her current level of alertness, and that the user must be able to correctly act upon this assessment.

Methods also exist to predict alertness level based on user inputs known empirically to modify alertness. U.S. Patent No. 5,433,223 to M. Moore-Ede et al. describes a method for predicting the likely alertness level of an individual at a specific point in time (past, current or future) based upon a mathematical computation of a variety of factors (referred to as "real-world" factors) that bear some relationship to alterations in alertness. The individual's Baseline Alertness Curve (BAC) is first determined based on five inputs and represents the optimal alertness curve displayed in a stable environment. Next, the BAC is modified by alertness modifying stimuli to arrive at a Modified Baseline Alertness Curve. Thus, the method is a means for predicting an individual's alertness level, not cognitive performance.

Another method has been designed to predict "work-related fatigue" as a function of number of hours on duty. Fletcher and Dawson describe their method in an article entitled "A Predictive Model of Work-Related Fatigue Based on Hours of Work" published in *Journal of Occupational Health and Safety*, vol. 13, 471-485 (1997). In this model a simplifying assumption is made - it is assumed that length of on-duty time correlates positively with time awake. To implement the method, the user inputs a real or hypothetical on-duty/off-duty (work/rest) schedule. Output from the model is a score that indicates "work-related fatigue." Although this "work-related fatigue" score has been shown to correlate with some performance measures, it is not a direct measure of cognitive performance per se. It can be appreciated that the fatigue score will be less accurate under circumstances when the presumed relationship between on-duty time and time awake breaks down - for example when a person works a short shift but then spends time working on projects at home rather than sleeping or when a person works long shifts but conscientiously sleeps all the available time at home. Also, this method is obtrusive in that the user must input on-duty/off-duty information rather than such information being automatically extracted from an unobtrusive recording device. In addition, the model is limited to predictions of "fatigue" based on work hours. Overall, this model is limited to work-related situations in which shift length consistently correlates (inversely) with sleep length.

Given the importance of the amount of sleep and the time of day for determining cognitive performance (and hence estimating productivity or effectiveness), and given the ever-increasing requirements of most occupations on cognitive performance, it is desirable to design a reliable and accurate method of predicting cognitive performance. It can be appreciated that increasing the number of relevant inputs increases cognitive performance prediction accuracy. However, the relative benefits gained from such inputs must be weighed against the additional burdens/costs associated with their collection and input. For example, although certain fragrances have been shown to have alertness-enhancing properties, these effects are inconsistent and negligible compared to the robust effects of the individual's sleep/wake history and the time of day. More important, the effect of fragrances on cognitive performance is unknown. Requiring an individual to keep a log of exposure to fragrances would be time consuming to the individual and only result in negligible gains in cognitive performance prediction accuracy. In addition, while the effects of the sleep/wake history and the time of day on cognitive performance are well known, the effects of other putative alertness-altering factors (e.g., job stress and workload), how to measure them (their operational definition), and their direction of action (cognitive performance enhancing or degrading) are virtually unknown.

An important and critical distinction between the present invention and the prior art is that the present invention is a model to predict performance on tasks with a cognitive component. In contrast, previous models involving sleep and/or circadian rhythms (approximately 24-hour) focused on the prediction of "alertness" or "sleepiness." The latter are concepts that specifically relate to the propensity to initiate sleep, not the ability to perform a cognitive task.

Although sleepiness (or its converse, alertness) could be viewed as an intervening variable that can mediate cognitive performance, the scientific literature clearly shows that cognitive performance and alertness are conceptually distinct, as reviewed by Johns in the article entitled, "Rethinking the Assessment of Sleepiness" published in *Sleep Medicine Reviews*, vol. 2, pp. 3-15 (1998), and as reviewed by Mitler et al. in the article entitled, "Methods of Testing for Sleepiness" published in *Behavioral Medicine*, vol. 21, pp. 171-183 (1996). Thomas et al. in the article entitled "Regional Cerebral Metabolic Effects of Prolonged Sleep Deprivation" published in *NeuroImage*, vol. 7, p. S130 (1998) reveal that 1-3 days



of sleep loss result in reductions in global brain activation of approximately 6%, as measured by regional cerebral glucose uptake. However, those regions (heteromodal association cortices) that mediate the highest order cognitive functions (including but not limited to attention, vigilance, situational awareness, planning, judgment, and decision making) are selectively deactivated by sleep loss to a much greater extent -- up to 50% -- after three days of sleep loss. Thus, decreases in neurobiological functioning during sleep restriction/deprivation are directly reflected in cognitive performance degradation. These findings are consistent with studies demonstrating that tasks requiring higher-order cognitive functions, especially those tasks requiring attention, planning, etc. (abilities mediated by heteromodal association areas) are especially sensitive to sleep loss. On the other hand, brain regions such as primary sensory regions, are deactivated to a lesser degree. Concomitantly, performance (e.g., vision, hearing, strength and endurance tasks) that is dependent on these regions is virtually unaffected by sleep loss.

Consequently, devices or inventions that predict "alertness" *per se* (e.g., Moore-Ede et al.) putatively quantify the brain's underlying propensity to initiate sleep at any given point in time. That is, devices or inventions that predict "alertness" (or its converse "sleepiness") predict the extent to which sleep onset is likely. The present invention differs from such approaches in that the nature of the task is accounted for -- i.e., it is not the propensity to initiate sleep that is predicted. Rather, the present invention predicts the extent to which performance of a particular task will be impaired by virtue of its reliance upon brain areas most affected by sleep deprivation (heteromodal association areas of the brain). The most desirable method will produce a highly reliable and accurate cognitive performance estimate based on the sleep/wake history of an individual and the time of day.

### III. DISCLOSURE OF THE INVENTION

A feature of the present invention is that it provides a numerical representation of predicted cognitive performance with an immediate ergonomic and economic advantage, i.e., an indication of productivity or effectiveness of an individual. Another feature of the present invention is that it does not require or use measurements/computations that are indirect, intermediate, inferential or hypothetical concomitants of cognitive performance. Examples of the latter are

alertness, sleepiness, time to sleep onset, body temperature and/or other physiological measures that vary with time. A further feature of the invention is that it accounts for transient or adventitious variations in cognitive performance from any source as a result of how that source affects the sleep/wake history (e.g., age) and/or physiological time of day (e.g., shift work). In effect, such sources are not treated as having effects on cognitive performance independent of the sleep/wake history and/or the time of day, and as such do not require separate measurement, tabulation, and input into the method.

One objective of this invention is to provide an accurate method for predicting cognitive performance of an individual.

A further objective is to provide a method that facilitates prediction of the effects of possible future sleep/wake histories on cognitive performance (forward prediction).

Another objective is to provide a method that facilitates retrospective analysis of likely prior cognitive performance based on the individual's sleep/wake history and the time of day.

Another objective is to provide a method for coordination and optimization of available sleep/wake time in order to obtain net optimal predicted cognitive performance for an individual and/or a group of individuals.

It can be appreciated that an implicit advantage and novelty of the method is its parsimony. The method uses those factors possessing maximal predictive value (as demonstrated empirically) as continuously updated inputs. Thus, the model will be simple to implement. Other models predicting "alertness" require the user to track multiple input variables (e.g., caffeine, alcohol ingestion, light/dark exposure, diurnal type), rather than presenting these inputs as optional "attachments" to a standard, simplified model based on those factors accounting for maximum cognitive performance change. For example, in accordance with a segment of the present method, the effects of age on cognitive performance are accounted for implicitly via the empirically derived effects of age on sleep. That is, sleep quality degrades with age. The inherent degradation in sleep quality with aging would implicitly result in a prediction of degraded cognitive performance (since in the present method degraded sleep results in a prediction of degraded cognitive performance), even if an individual's age were unknown. Therefore, age

need not constitute a separate (independent) input variable to a cognitive performance prediction model.

The invention also provides other significant advantages. For example, an advantage of this invention is the elimination of a need for empirical evaluation.

5 Another advantage of this invention is obtaining an accurate prediction of cognitive performance of an individual. The advantage may be achieved by a method incorporating two factors that have been empirically demonstrated to exert a significant effect on cognitive performance, namely, (1) the individual's sleep/wake history and (2) the time of day ("day" herein referring to a 24-hour  
10 period including both nighttime and daylight hours).

Another advantage achieved by this invention is an accurate prediction of current cognitive performance.

Another advantage achieved by this invention is that it is capable of providing a real time prediction of cognitive performance.

15 Yet another advantage achieved by this invention is a prediction of future expected cognitive performance throughout the day based on hypothetical future sleep/wake periods.

An additional advantage achieved by this invention is a retrospective analysis of cognitive performance at given times.

20 Another advantage of the invention is that a particular cognitive performance prediction is not based on normative data (i.e., does not require a "look-up table" for output), but rather is calculated directly based on each individual's sleep/wake information and the time of day.

Another advantage of the invention is that it can be used to optimize the  
25 individual's future sleep/wake schedule based on a fixed mission/work schedule. Previous methods and apparatuses are directed toward modifying the work schedule and/or mission to "fit the individual". In most situations, however, work schedules and/or missions are fixed. Thus, modifying the work schedule or mission to suit the individual is impractical or impossible. A more reasonable  
30 approach incorporated in the present method is to allow the individual to adjust his/her sleep/wake periods to meet work/mission demands. Thus, the current method presents a more practical alternative by providing a means to regulate work hours to a directly applicable metric (cognitive performance) instead of

regulating work hours by time off duty or by using indirect measures of cognitive performance such as alertness.

A feature of this invention is the provision of a graphical representation that translates an individual's sleep/wake history and the time of day into an immediately useful, self-evident index. A prediction of cognitive performance, unlike a prediction of "alertness" or "sleepiness", requires no further interpretation.

The system for predicting human cognitive performance based on information from an actigraph in accordance with the invention accomplishes the above objectives and achieves the above advantages. The system is easily adapted to a wide variety of situations and types of inputs.

In accordance with an aspect of the invention: an individual activity is monitored by an actigraph, the activity information is converted into an individual's sleep/wake history, which is inputted into a processing device. The processing device classifies the individual pieces of sleep/wake history data as either sleep or wake. Based on the classification of data, the processing device selects and calculates a cognitive performance capacity corresponding to the present state of the individual, the cognitive performance capacity may be modified by a time of day value to adjust the cognitive performance capacity to a predicted cognitive performance. The predicted cognitive performance represents the ability of the individual to perform cognitive tasks. The predicted cognitive performance may be displayed for a real-time indication or as part of a curve, printed out with the information that could have been displayed, and/or stored for later retrieval and/or use. The calculation of the cognitive performance capacity is made based on functions which model the effect of the interrelationship of sleep and being awake on cognitive performance. The time of day function models the effect of an individual's circadian rhythms on cognitive performance.

In accordance with the underlying method of the invention, the method can be accomplished with a wide variety of apparatus. Examples of the possible apparatus embodiments include electronic hardware as either a dedicated equipment or equipment internal to a computer, software embodied in computer readable material for use by computers, software resident in memory or a programmed chip for use in computers or dedicated equipment, or some combination of both hardware and software. The dedicated equipment may be part of a larger device that would complement the dedicated equipment's purpose.

Given the following enabling description of the drawings, the invention should become evident to a person of ordinary skill in the art.

#### IV. BRIEF DESCRIPTION OF THE DRAWINGS

Figure 1(a) is a block diagram representation of the modulation in the preferred embodiment. Figure 1(b) graphically shows the combination of output from the functions represented by Figure 3(a) with time of day modulation to derive predicted cognitive performance.

Figure 2 is a block diagram representation of the wake, sleep, delay, and sleep inertia functions for calculating predicted cognitive performance capacity.

Figure 3(a) graphically illustrates the effect of being awake and asleep on cognitive performance capacity over a 24-hour period. Figure 3(b) is an enlarged view of circled portion 3(b) of Figure 3(a), and graphically shows the delay function with respect to cognitive performance capacity. Figure 3(c) is an enlarged view of circled portion 3(c) of Figure 3(a), and graphically shows the sleep inertia function with respect to cognitive performance capacity.

Figures 4(a) and (b) depict a detailed flowchart showing the steps of the method of the invention.

Figure 5 illustrates a functional representation of an alternative embodiment.

Figure 6 depicts a block diagram of structural components for the preferred embodiment.

Figures 7(a) and (b) illustrate a detailed flowchart showing the steps of an alternative embodiment.

Figure 8 illustrates predicted cognitive performance with 8 hours of sleep per night.

Figure 9 illustrates predicted cognitive performance with 4 hours of sleep per night.

Figure 10 illustrates predicted cognitive performance with no sleep during 72 hours of total sleep deprivation.

Figure 11 illustrates predicted cognitive performance with a recovery sleep of 8 hours the night after total sleep deprivation.

Figure 12 illustrates predicted cognitive performance with daily 30-minute naps during 85 hours of sleep deprivation.

Figure 13 illustrates predicted cognitive performance with sleep fragmented with 10 arousals per hour.

Figure 14 illustrates predicted cognitive performance for two nights of shift work with daytime sleep.

Figure 15 illustrates predicted cognitive performance prior to a commercial vehicle driver collision.

5        Figure 16 illustrates predicted cognitive performance with 15 hours on-duty/8 hours off-duty schedule with 6 hours sleep per night during duty days.

Figure 17 illustrates predicted cognitive performance with alternative 12 hours on-duty/12 hours off-duty schedule with 8 hours sleep per night during duty days.

10       Figure 18(a) illustrates an actigraph circuit. Figure 18(b) illustrates an actual actigraph based on the Figure 18(a) actigraph circuit.

Figure 19(a) illustrates another actigraph circuit. Figure 19(b) illustrates a top view of an actual actigraph based on the Figure 19(a) actigraph circuit. Figure 19(c) illustrates a bottom view of an actual actigraph based on the Figure 19(a) actigraph circuit.

## **V.    MODES FOR CARRYING OUT THE DESCRIBED EMBODIMENTS**

The present invention involves a method for predicting cognitive performance at a given time in the past, present, or future as a consequence of the amount of sleep and wakefulness up to that time and further as a function of the time of day. The method calculates a numerical estimate of cognitive performance for an individual as a continuous function of time. The calculations (described below) are based on empirically derived direct mathematical relationships among (1) the continuous decrement of cognitive performance during wakefulness; (2) restoration of cognitive performance during sleep; and (3) cyclic variation in cognitive performance during the course of the day.

25       In accordance with the invention, a numeric value indicating predicted cognitive performance at a given moment in time is provided as shown in Figures 1(a)-4(b). As shown in Figures 1(a)-(b), predicted cognitive performance equals the output of a series of calculations obtained in three general steps, using functions empirically derived from direct measurements of cognitive performance under scientifically controlled conditions. The first step, as shown in Figure 2, uses a set of functions to calculate an initial value referred to as the level of cognitive performance capacity as graphically depicted in Figures 3(a)-(c). Once the level of cognitive performance capacity is calculated, the second step calculates or uses a

previously calculated time of day modulator  $M$  represented as G8 in Figure 1(b) and S8 in Figure 4(b). The third step involves the mathematical combination of the results from the first and second steps yielding predicted cognitive performance, shown as a block diagram in Figure 1(a) and graphically represented in Figure 1(b).

There are four functions relating to the sleep/wake history used to calculate the level of cognitive performance capacity as shown in Figures 2-4(b). The wake function  $w(t)$  quantifies empirically derived relationships between the time awake and degradation of cognitive performance. The sleep function  $s(t)$  quantifies empirically derived relationships between the time asleep and maintenance and/or recuperation of cognitive performance. In addition to these two primary functions that operate during the bulk of the time awake or asleep there are two other functions that operate briefly during the transition from one state to the other. They include the delay of recuperation function  $d(t)$  and the sleep inertia function  $i(t)$ . The delay of recuperation function  $d(t)$  represents the relationship between the wake to sleep transition and the recuperation of cognitive performance. This function operates during the initial period of sleep following being awake as shown in Figure 3(b). The sleep inertia function  $i(t)$  represents the relationship between the sleep to wake transition and cognitive performance. This function operates during the initial period of time being awake after being asleep as shown in Figure 3(c).

The function representing the time of day  $M$ 's effects on cognitive performance is used to calculate a modulating factor. The time of day function describes empirically derived relationships between the time of day (point in time within a 24-hour period) and the variation in cognitive performance over the course of the day as exemplified by G8 in Figure 1(b).

A mathematical operation, shown in Figure 1(b) as multiplication, is used to combine the results from the first and second steps into a single predicted cognitive performance curve  $E$  in the third step.

The inputted data S2 into the method includes a representation of an individual's sleep/wake history. The sleep/wake history is a time series or temporal record based on local clock time. Each successive period, interval or epoch identifies one of two mutually exclusive states, namely wake or sleep. Such a sleep/wake history is not necessarily "historical" in the sense of occurring in the

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software to analyze the data recorded by the actigraph including sleep scoring algorithms.

More recently, digital signal processing (DSP) actigraphs have begun to be developed. Because the DSP actigraph will provide much more information than just  
5 the conventional number of zero crossings or counts above threshold (this and other information provided by a conventional actigraph will, however, be retained), it shows promise for distinguishing between different sleep stages. Thus, sleep scoring systems for DSP will not only replace, but will also make irrelevant, the Cole-Kripke algorithm. A sleep scoring system for the DSP will be developed as the DSP  
10 database of empirical data from use of DSP actigraphs increases.

Other algorithms and methodologies for automated actigraphy scoring have been developed by, for example, Jean-Louis et al., 1996; Sadeh et al., 1989; and Zisapel et al., 1995. Each of these scoring systems shows considerable promise, especially for scoring the actigraphically recorded sleep/wake states of individuals  
15 with sleep disorders or other medical disorders. Available scoring systems mainly differ along technical aspects, for example, the extent to which activity counts in previous and subsequent epochs influence the scoring of the current epoch; and variation among mathematical principles underlying each scoring system. As one of ordinary skill in the art will realize from reading this description, any actigraph  
20 scoring system is capable of providing the sleep/wake data input for the method of this invention.

The sleep/wake history will preferably take the form of a data series. The sleep/wake history may include past, present, and/or future (predicted) sleep/wake patterns of an individual. The sleep/wake history is a representation of a state of  
25 an individual as either being asleep or awake and is divided into epochs. The epochs are the same length, but that length could be of any time period as dictated by restraints of the method and apparatus used to collect data and/or the desired precision of the sleep/wake pattern.

It can be appreciated that the accuracy of the cognitive performance  
30 prediction is directly related to the accuracy of the sleep/wake history input and the sleep scoring system used to interpret the sleep/wake states of an individual. One possible source of inaccuracy may arise from the temporal resolution of the input epoch or interval. That is, the shorter the input epoch, the higher the temporal resolution and consequent moment-to-moment accuracy of the sleep/wake input.

For example with actigraphy, past experience indicates that the most effective length of an epoch is one minute. Another source of inaccuracy may arise from ambiguity in the sleep/wake discrimination itself. In the event that the history input is ambiguous (i.e., the sleep or wake state is uncertain), the calculation of predicted cognitive performance can be performed twice concurrently, once for each possible state (sleep or wake), resulting in a dual output representing the possible range of expected cognitive performance. One of ordinary skill in the art will appreciate that the dual output can be further divided if there is more than one ambiguity in the sleep/wake history. Such treatment in executing the functions expressed below is included as a component of this method and any implementing apparatus.

The method of this invention is not limited with regard to time or technique: on-line/real-time vs. off-line/post-hoc; or incremental, iterative vs. discrete solutions to continuous forms of those equations.

A preferred embodiment of the method encompasses a mathematical model that expresses predicted cognitive performance capacity  $E$  at time  $t$  as a modulation of the current cognitive performance capacity  $C$  by a time of day function  $M$ . It can be written as a general description in its simplest form as:

$$E = C \nabla M \quad \text{Equation 1}$$

where  $\nabla$  represents a mathematical operator. Any mathematical operator may be used to combine cognitive performance capacity  $C$  and day of time function  $M$ . The form and nature of time of day function  $M$  dictates the exact operator that is most desirable. Most preferably, Equation 1a below would be used to combine cognitive performance capacity  $C$  and day of time function  $M$ .

$$E = C * M \quad \text{Equation 1a}$$

In the alternative, Equation 1b below could also be used to combine cognitive performance capacity  $C$  and day of time function  $M$ .

$$E = C + M \quad \text{Equation 1b}$$

Cognitive performance capacity  $C$  represents a function of sleep/wake history, that is

$$C = w(t) + s(t) + d(t) + i(t) \quad \text{Equation 2}$$

where  $w(t)$ ,  $s(t)$ ,  $d(t)$ , and  $i(t)$  are the instantaneous values of the wake, sleep, delay, and sleep inertia functions at time  $t$ . Time of day function  $M$  represents a function of the time of day, such that

$$M = m(t) \quad \text{Equation 3}$$

where  $m(t)$  is the instantaneous value of the time of day function at time  $t$ .

In keeping with the invention, a three-step process may be performed after either an initial setting of the starting time  $t$ , the starting cognitive performance capacity  $C$ , and the time of the last transition  $t_{LS}$  when appropriate in S1 of Figure 4 where these data can be entered in any order. In the first step, the level of cognitive performance capacity  $C$  at time  $t$  may be calculated based on an individual's sleep/wake history using functions  $w(t)$ ,  $s(t)$ ,  $d(t)$ , and  $i(t)$  as represented by S3-S7e in Figures 4(a) and (b). In the second step, time of day modulator  $M$  may be calculated using the time of day function as represented by S8 in Figure 4. According to an aspect of the invention, the second step can be performed once to provide a series of data points in time sequential order for multiple executions of the first step. In the third step, predicted cognitive performance  $E$  may be derived from the combination of cognitive performance capacity  $C$  and time of day modulator  $M$  resulting in cognitive performance capacity  $C$  being modulated by time of day modulator  $M$  as illustrated by S9 in Figure 4(b).

#### **First Step: Calculation of Cognitive Performance Capacity $C$**

Figure 2 is a schematic flow diagram representing the use of the functions described below. Examples of the calculations discussed are graphically illustrated in Figures 3(a)-(c). Figures 4(a) and (b) are a detailed flowchart of the steps in the method. As a preferred embodiment of the model, cognitive performance capacity  $C$  is herein assigned values having a total range of zero to 100 and thus represent percentages. The ranges in this application are intended to encompass the end points of the stated numerical range. However, cognitive performance capacity  $C$  may be scaled to other values or units for specific applications.

In the preferred embodiment, only one of the four functions  $w(t)$ ,  $s(t)$ ,  $d(t)$ , and  $i(t)$  operates at any given interval of time, and the others are equivalent to zero in Equation 2 as represented by S7a through S7d. Functions  $w(t)$  and  $s(t)$  describe the non-transition states, while functions  $d(t)$  and  $i(t)$  describe the transition states. For instance in a non-transition state, when the individual is awake, function  $s(t)$  is set to zero, and when the individual is asleep, function  $w(t)$  is set to zero. Likewise, during specific intervals of transition from wake to sleep and vice versa, only one of the transition functions  $d(t)$  or  $i(t)$  operates, the other being set equal to zero. When there is a change between sleep and wake, or vice versa, a time counter  $t_{LS}$

is reset to keep track of the time in the present state for determining decision rules for the transition functions  $d(t)$  and  $i(t)$  as shown in Figure 4(b).

(1) Wake function ( $w(t)$ )

The wake function S7a represents the depletion of cognitive performance capacity with the passage of time awake. It is based on evidence that (1) near-100% cognitive performance is maintained from day to day when individuals obtain eight hours of sleep each night; and (2) cognitive performance appears to decline by approximately 25% for every 24 hours of wakefulness.

In S7a, the wake function  $w(t)$  calculates the current value of cognitive performance capacity  $C$  resulting from the decay in cognitive performance capacity that occurs over an interval of time from  $t-1$  to  $t$ , which in the preferred embodiment is the length of one epoch. As noted above, this calculation is performed independent of and prior to modulation of cognitive performance capacity  $C$  by the time of day function  $M$  in S9. A generalized form of the wake function is given by the equation:

$$C_w = w(t)$$

Equation 4

where wake function  $w(t)$  may be any positive-valued function decreasing with  $t$ . More preferably, the wake function  $w(t)$  is a linear function depleting performance at a constant rate, and, most preferably, the wake function  $w(t)$  is expressed at time  $t$  as follows:

$$w(t) = C_{t-1} - k_w$$

Equation 4a

where the interval of wakefulness is from  $t-1$  to  $t$  (in epochs) and the decay in performance per minute is  $k_w$ . Thus, if  $t-1$  to  $t$  is not one minute, then  $k_w$  is appropriately adjusted. The total range of  $k_w$  is any positive real number, and preferably  $k_w$  is a range of .003 to .03% per minute, and most preferably  $k_w$  is equal to approximately 1% per hour or 0.017% per minute. The value  $k_w$  is based on empirical data showing that cognitive performance declines by approximately 25% for every 24 hours of continuous wakefulness. Equation 4a is represented in Figures 2 and 4(b) at S7a. An example is illustrated as the wake function in Figure 3(a), for an initial cognitive performance capacity of 100%, a decay rate of 0.017% per minute, over an interval of 16 hours (960 minutes).

(2) Sleep function ( $s(t)$ )

The sleep function S7c restores cognitive performance capacity with the passage of time asleep. The sleep function  $s(t)$  is based on empirical evidence

that the recuperative value of sleep on cognitive performance accumulates in a nonlinear manner. That is, the rate of cognitive performance capacity recuperation is higher initially during sleep and slows as the time asleep accumulates. Other data indicates that sleep beyond a certain point confers little or no additional benefit for cognitive performance and the rate of recuperation approaches zero. Thus, for example, two hours of sleep are not twice as recuperative as one hour of sleep. The sleep function increases cognitive performance capacity at a rate that depends on the current level of cognitive performance capacity -- the lower the initial cognitive performance capacity, the more rapidly recuperation accumulates. For example, following a full day (16 hours) of wakefulness, during ensuing nighttime sleep recuperation accumulates rapidly early in the night. As cognitive performance capacity is restored across the sleep period, the rate of recuperation declines. Following sleep deprivation, initial cognitive performance capacity is even lower than it would be following a normal 16-hour day, and the rate of recuperation is even higher than at the beginning of recovery sleep. During chronic partial sleep deprivation, cognitive performance capacity may not be completely restored each night despite this more rapid initial recuperation rate.

The sleep function calculates the current value of cognitive performance capacity  $C$  resulting from the recovery of capacity that occurs while an individual is asleep over an interval of time  $T$  (from  $t-1$  to  $t$ ). As noted above, this calculation is performed independent of, and prior to, modulation of  $C$  by the time of day function  $M$ . A generalized form of the sleep function is given by the equation:

$$C_s = s(t) \quad \text{Equation 5}$$

where sleep function  $s(t)$  may be any positive-valued function increasing with  $t$ , and more preferably the sleep function  $s(t)$  is an exponential function. This is based on empirical data showing that cognitive performance restoration during sleep is nonlinear, with the rate of recuperation highest initially and gradually slowing as sleep continues. Thus, the most preferred sleep function is an exponential function, which in its discrete form is stated as:

$$C_t = C_{t-1} + (100 - C_{t-1}) / k_s \quad \text{Equation 5a}$$

where the interval of sleep is from  $t-1$  to  $t$  (in minutes), the maximum cognitive performance capacity value is 100%,  $C_{t-1}$  is cognitive performance capacity in the period preceding time  $t$ , and  $k_s$  is the recuperation "time constant". In other words,  $k_s$  is the time required to fully restore cognitive performance capacity  $C$  if it was

restored at a constant rate equal to the initial slope of the curve. The recuperation time constant  $k_s$  is derived empirically from partial sleep deprivation data and is selected based on the length of the epoch. In accordance with the preferred embodiment,  $k_s$  is equal to any positive real number. For example,  $k_s$  may be in the range of 100 to 1000, and, more particularly may be approximately 300 with an epoch length of one minute. However, the optimum values for  $k_s$  will depend at least in part on the length of the epoch. Equation 5a is represented in Figures 2 and 4(b) as S7c. A graphical example is illustrated as the sleep function in Figure 3(a), using an initial cognitive performance capacity level of 100, and using a time period of one minute and  $k_s = 300$ , the effect of eight hours of sleep following 16 hours of wakefulness.

(3) Delay function  $d(t)$  for wake to sleep transitions

The delay of recuperation function  $d(t)$  defines the duration of an interval after sleep onset during which recuperation of cognitive performance capacity from the sleep function is delayed. During this interval, the wake function degradation of cognitive performance capacity continues as represented by S7d in Figure 4(b). By preventing immediate accumulation of cognitive performance capacity at the beginning of a sleep period or following awakenings from sleep, this delay adjusts the cognitive performance capacity calculation S6b.

The delay of recuperation function is based upon empirical studies showing that the first few minutes of sleep are generally comprised of stage 1 sleep, which is not recuperative for sustaining cognitive performance capacity. Frequent arousals to wake or stage 1 sleep (sleep fragmentation) drastically reduce the recuperative value of sleep on cognitive performance capacity. Available data suggest that five minutes is the approximate length of time required to return to recuperative sleep (stage 2 or deeper sleep) following an arousal to wake or stage 1 sleep. If many hours of sleep are obtained without interruption, then the delays make only a small difference in overall restoration of cognitive performance capacity. If sleep is interrupted with frequent awakenings, the delays in recuperation after each awakening will accumulate, and thus substantially reduce total cognitive performance capacity restored during the total sleep period.

The delay function specifies the duration of a sleep interval during which application of the sleep function is postponed and a transitional formula is applied.

A generalized form of the delay function for wake to sleep transitions is expressed as a decision rule:

$$\begin{array}{ll}
 d(t): & \text{IF } (t - t_{LS}) \leq k_d \\
 & \text{THEN } C_t = d(t) \\
 5 & \text{ELSE } C_t = s(t)
 \end{array}
 \qquad \text{Equation 6}$$

where LS stands for last state change, and thus the wake to sleep transition time  $t_{LS}$  denotes the time of the last wake interval preceding a contiguous series of sleep intervals. This decision rule is shown in Figures 2 and 4(b) as S6b, S7c and S7d taken together. For calculating cognitive performance capacity during the interval  $k_d$ , cognitive performance capacity  $C_t$  is evaluated by a transitional formula  $C_t = d(t)$ . After  $k_d$  has elapsed,  $C_t = s(t)$ . Note that if wakefulness ensues before the end of  $k_d$ , then  $C_t$  never reverts to  $s(t)$ . That is the sleep function is not applied during the brief sleep interval.

It is believed that the preferred range for  $k_d$  is from 0 to 30 minutes, more preferably  $k_d$  equals about five minutes measured from the time of sleep onset before recuperation is derived from sleep. Preferably  $d(t)$  equals  $w(t)$ . One of ordinary skill in the art will realize there are a variety of factors that influence the length of  $k_d$ . Thus a more preferred delay function may be expressed as:

$$\begin{array}{ll}
 d(t): & \text{IF } (t - t_{LS}) \leq 5 \\
 20 & \text{THEN } C_t = w(t) \\
 & \text{ELSE } C_t = s(t)
 \end{array}
 \qquad \text{Equation 6a}$$

The effects of delayed recovery on cognitive performance capacity, as embodied by Equation 6a, are graphically illustrated in detail in Figure 3(b).

#### (4) Sleep inertia function $i(t)$ for sleep to wake transitions

25 The sleep inertia function  $i(t)$  defines the duration of an interval after awakening from sleep during which manifest cognitive performance capacity is suppressed below the actual current level. The sleep inertia function  $i(t)$  is based upon empirical data showing that cognitive performance is impaired immediately upon awakening, but improves primarily as a function of time awake. It is also  
30 based on positron emission tomography studies showing deactivated heteromodal association cortices (those areas that mediate this cognitive performance) immediately upon awakening from sleep, followed by reactivation of these areas over the ensuing minutes of wakefulness. That is, actual cognitive performance recuperation realized during sleep is not apparent immediately after awakening.

The data indicate that 20 minutes is the approximate length of time required for cognitive performance capacity to return to levels that reflect actual recuperation accrued during sleep.

A sleep inertia delay value  $k_i$  specifies the duration of the interval after  
 5 awakening during which manifest cognitive performance capacity may be transitionally suppressed below the sleep-restored cognitive performance capacity level. During this interval, a transitional function bridges from an initial level to that determined by the wake function alone. A generalized form of the sleep inertia function for sleep to wake transitions is expressed as a decision rule:

10  $i(t)$ : IF  $(t - t_{LS}) < k_i$   
           THEN  $C_t = i(t)$   
           ELSE  $C_t = w(t)$

Equation 7

where the sleep to wake transition time  $t_{LS}$  denotes the time of the last sleep interval preceding a contiguous series of wake intervals. For calculating cognitive  
 15 performance capacity during the interval  $k_i$ ,  $C_t$  is evaluated by a transitional formula  $C_t = i(t)$ . After  $k_i$  has elapsed,  $C_t = w(t)$ . Equation 7 is represented in Figures 2 and 4(b) as S6a, S7a and S7b taken together.

The preferred range for  $k_i$  is from 0 to about 60 minutes, and preferably in the range of about 10 to about 25 minutes, and most preferably between 18 and 22  
 20 minutes.

The sleep inertia function  $i(t)$  may be any function over the interval 0 to  $k_i$ , preferably any negatively accelerated function. A preferred sleep inertia function  $i(t)$  is a simple quadratic equation. This function preferably suppresses cognitive performance capacity by 10% to 25% immediately upon awakening, and most  
 25 preferably by 25%. The function recovers 75% of the suppressed cognitive performance capacity in the first 10 minutes after awakening and 100% of the suppressed cognitive performance capacity usually by 20 minutes after awakening, after which the wake function resumes. These values are based on empirical data concerning the transition from sleep to wake. These studies show that cognitive  
 30 performance is impaired immediately upon awakening from sleep, that the bulk of this impairment dissipates within the first few minutes of awakening, and that approximately 20 minutes is required for performance to be fully restored. Using the preferred 25% suppression of cognitive performance capacity and 20 minute



recovery time, the preferred form of the sleep inertia function is expressed as a decision rule:

$$\begin{aligned}
 i(t): \quad & \text{IF } (t - t_{LS}) < 20 \\
 & \text{THEN } C_t = C_{SW} * [0.75 + 0.025(t - t_{LS}) - (0.025(t - t_{LS}))^2] \\
 & \text{ELSE } C_t = w(t)
 \end{aligned}$$

Equation 7a

where  $C_{SW}$  is cognitive performance capacity at the end of the sleep period calculated by the sleep function at the sleep to wake transition time  $t_{LS}$ . This decision rule is shown in Figures 2 and 4 as S6a, S7a, and S7b taken together. Equation 7a illustrates an initial suppression of 25% and  $k_i$  equal to 20 minutes, and a negatively accelerated ramp bridging the interval until the wake function  $w(t)$  resumes its effects. The effect of the sleep inertia function  $i(t)$  on cognitive performance capacity, as embodied by Equation 7a, is graphically illustrated in Figure 3(c).

An alternative variant of the sleep inertia function  $i(t)$  is a linear equation based on  $k_i$  equal to 10 minutes and an initial 10% decrease in cognitive performance capacity. The resulting decision rule is then:

$$\begin{aligned}
 i(t): \quad & \text{IF } (t - t_{LS}) < 10 \\
 & \text{THEN } C_t = C_{SW} * [0.9 + (t - t_{LS}) / 100] \\
 & \text{ELSE } C_t = w(t)
 \end{aligned}$$

Equation 7b

As one of ordinary skill in the art will realize, both Equations 7a and 7b can be adjusted for a change in the value of  $k_i$  and the initial suppression of cognitive performance capacity.

Second Step: Calculation of the time of day modifier M

(1) Time of day function m(t)

The time of day function  $m(t)$  shown at S8 in Figure 4(b) describes the cyclical 24-hour variation in cognitive performance. The time of day function  $m(t)$  is based on empirical data showing that under constant routine and/or total sleep deprivation conditions (i.e., with sleep/wake history controlled), cognitive performance capability oscillates between approximately 5% to approximately 20% peak to peak over a 24-hour period. This effect is commonly attributed to circadian rhythms of individuals. Output from this function modulates the current cognitive performance capacity prediction C (calculated in the first step) according to the time of day. The result of this modulation is the predicted cognitive performance capacity E. A generalized form of the time of day function is given by

$$M = m(t)$$

Equation 8

where  $m(t)$  can be any rhythmic function with a base period of 24 hours, and, preferably,  $m(t)$  is the sum of two sinusoids, one with a period of 24 hours and the second with a period of 12 hours, which provides a biphasic circadian component.

5 This function may be based on empirical data showing that a considerable proportion of variability seen in cognitive performance measurements can be accounted for by two such sinusoidal waveforms. As previously noted, the peak in empirically observed cognitive performance capacity occurs usually between 8:00 PM and 10:00 PM, and the trough occurs usually between 2:00 AM and 6:00 AM,  
10 providing a variation of approximately 5% to approximately 20% each day. A secondary trough occurs usually around 3:00 PM. Using these values for the preferred form of function  $m(t)$ , the resulting function accounts for the empirically demonstrated asymmetry of daily cognitive performance rhythms, with a mid-afternoon decrease.

15 The descriptive form of the function  $m(t)$ , including its offset and amplitude values varies with the operator selected for the third step. The computed value of the function can be expressed either as an additive percentage of cognitive capacity (dependent or independent of the current value of cognitive performance capacity  $C_t$ ) or as a multiplicative dimensionless scalar. The preferred form of the  
20 function, using the multiplicative operator, is expressed as

$$m(t) = F + (A_1 * \cos(2\pi(t - V_1) / P_1) + A_2 * \cos(2\pi(t - V_2) / P_2))$$

Equation 8a

where  $F$  is an offset,  $t$  is the time of day,  $P_1$  and  $P_2$  are periods of two sinusoids,  $V_1$  and  $V_2$  are the peak times of day in time units or epochs past midnight, and  $A_1$  and  
25  $A_2$  are amplitudes of their respective cosine curves. This function may be used to modulate the previously calculated cognitive performance capacity  $C$ , the result being the predicted cognitive performance capacity  $E$ . Equation 8a is shown as S8 in Figures 1(a) and 4(b) and graphically illustrated as G8 in Figure 1(b). As shown in Figure 4(b),  $t$  is an input in the time of day function  $m(t)$  for each epoch of data.

30 For example in a preferred embodiment the variables are set as follows:  $t$  is the number of minutes past midnight,  $P_1$  is equal to 1440 minutes,  $P_2$  is equal to 720 minutes,  $V_1$  is equal to 1225, and  $V_2$  is equal to 560. Further, when  $A_1$  and  $A_2$  are represented as scalars, their amplitudes are in a range from 0 to 1, and more preferably are in a range from 0.01 to 0.2, and most preferably  $A_1$  is equal to 0.082

and  $A_2$  is equal to 0.036. Further in this example  $F$  is equal to either 0 or 1, and more preferably  $F$  is equal to 1. The resulting value of the time of day function  $m(t)$ , in this example, is in the range of 0 to 2, and preferably in the range of 0.8 to 1.2, and most preferably in the range of 0.92 to 1.12.

### 5 Third Step: Calculation of Predicted Cognitive Performance

The overall process of calculating predicted cognitive performance capacity  $E$  is illustrated schematically in Figures 1(a)-(b) and 4(a)-(b). The time of day function  $M$  modulates the cognitive performance capacity  $C$  derived from the individual's sleep/wake history to generate the final predicted cognitive  
 10 performance  $E$  as shown in Figures 1(a)-(b). In the third step, predicted cognitive performance  $E$  is derived from the combination of cognitive performance capacity  $C$  and time of day function  $M$ . In its most general form:

$$E = C \nabla M$$

Equation 1

where  $\nabla$  is any mathematical operation for combining cognitive performance  
 15 capacity  $C$  and time of day function  $M$ . The conventional choice of operations for providing this combination is addition or multiplication. Depending on the form of time of day function  $m(t)$  selected above, the same numerical value of predicted cognitive performance  $E$  can be generated by either operation. Most preferably the combination is performed with multiplication S9, represented as:

$$20 \quad E = C * M$$

Equation 1a

In Equation 1a, the predicted cognitive performance  $E$  is the modulation of the current cognitive performance capacity  $C$  and a value centered around the number one representing the current value of the time of day modulator  $M$ .

As noted above, the preferred numerical representation of cognitive  
 25 performance capacity  $C$  is a value ranging from zero to 100 to represent a percentage of cognitive performance capacity available. However, predicted cognitive capacity  $E$  can meaningfully exceed 100 under certain circumstances due to time of day modulation about the current value of cognitive performance capacity  $C$ . A possible example of such a circumstance would be a sleep period resulting in  
 30 100% cognitive performance capacity  $C$  and terminated at the evening peak (and after sleep inertia has dissipated). To retain the 100% scale, either the predicted cognitive capacity  $E$  may be truncated/clipped at 100% or 0 to 120% may be scaled to 0 to 100%. Either choice will maintain a maximum of 100%. This most

likely will be implemented as scaling 120% to 100% and then truncating/clipping any predicted cognitive capacity  $E$  to 100%.

As shown in Figure 1, the method repeats for each new epoch of data. For each iteration of the method, one time unit equal to the length of an epoch may be added to time  $t$  preferably in the form of a counter  $S11$  as exemplified in Figure 1.

In the preferred embodiment described above, the sleep inertia function  $i(t)$  is applied to cognitive performance capacity  $C$  prior to modulation of cognitive performance capacity  $C$  by the time of day modifier  $M$ . An alternative embodiment applies the sleep inertia function  $i(t)$  not to cognitive performance capacity  $C$ , but to predicted cognitive capacity  $E$ , that is, subsequent to the modulation of cognitive performance capacity  $C$  by time of day modifier  $M$ . The state of empirical knowledge is insufficient to determine whether the preferred embodiment is better than this alternative embodiment.

Also in the preferred embodiment described above, the wake function  $w(t)$  is set to zero when the sleep inertia function  $i(t)$  is applied. Another alternative embodiment applies the sleep inertia function  $i(t)$  and the wake function  $w(t)$  simultaneously. When the sleep inertia function  $i(t)$  and the wake function  $w(t)$  become equal to each other or the sleep inertia function  $i(t)$  becomes greater than the wake function  $w(t)$ , then cognitive performance capacity  $C$  is calculated using the wake function  $w(t)$ .

The preferred embodiment may be further modified to account for the effects of narcotics or other influences that will impact the cognitive capacity as shown in Figure 5. Further modification to the preferred embodiment will allow for the inclusion of jet lag and similar time shifting events by, for example, compressing or expanding the 24 hour period of the time of day function  $M(t)$  over a period of days to realigning the time of day function  $M(t)$  to the adjusted schedule.

#### **Implementation of the Method**

The preferred embodiment may be realized as software to provide a real-time current state of an individual's cognitive performance and the capability upon demand to extrapolate future levels of cognitive performance. A flowchart representing the steps to be performed by the software in the preferred embodiment is shown in Figures 4(a)-(b) and for an alternative embodiment, to be described later, in Figures 7(a)-(b). The software may be implemented as a computer program or other electronic device control program or an operating

system. The software may be resident in an actigraph, if the actigraph also includes sleep scoring capabilities, attached to the individual or may be resident in a stand-alone device. Alternatively, the software may be resident in a stand-alone device in communication with the actigraph at intervals or continuously. The stand-alone device may be a personal computer, a PAL device, a personal digital assistant (PDA), an e-book or other handheld or wearable computing devices (incorporating Palm OS, Windows CE, EPOC, or future generations like code-named Razor from 3Com or Bluetooth from consortium including IBM and Intel), or a specific purpose device receiving signals from an actigraph attached to an individual. Depending on the location of the software, the software could be stored, for example, in random access memory (RAM); in read only memory (ROM); on a storage device like a hard drive, disk, compact disc, punch card, tape or in other computer readable material; in virtual memory on a network such as an intranet or the Internet, computer or otherwise; on an optical storage device; on a magnetic storage device; and/or on an EPROM. Alternatively, the software may allow for the variables in the equations discussed above to be adjusted and/or changed. This capability will allow users to adjust the variables based on empirical knowledge and also learn the interrelationship between the variables.

When the software is resident on the actigraph, the software will convert any decimal numbers used in calculations into integers that are appropriately scaled as is well known to those skilled in the art. Further the integers would then be approximated such that minimal error would be created, for example, approximation for the Cole-Kripke algorithm weighting factors become 256, 128, 128, 128, 512, 128, and 128, respectively. Using linear approximation will simplify the binary arithmetic and the corresponding assembly code for software implementation. Preferably, the time of day modulator would be embodied as a table with one hour steps resulting in 24 rows using 8-bit unsigned integers. The intervening steps would be interpolated from the one hour steps to provide 15-minute steps. This simplification provides sufficient resolution for available displays. As the resolution of available displays improves, smaller temporal steps may be used for the table and/or interpolation to replicate the time of day modulator. As the resolution of available displays improves, smaller temporal steps may be used for the table and/or interpolation to replicate the time of day modulator. A pointer system would be utilized to retrieve the appropriate data to

calculate the time of day modulator. Depending on a myriad of factors, one of ordinary skill in the art will most likely choose a multiplicative modulation to achieve appropriate scaling or an additive modulation for less complex but more rapid evaluation, i.e., if speed is a concern. The main disadvantage with the additive modulation is that there will be an approximately 3% error compared to the 1% error using the multiplicative modulation in this invention. This system will allow the time of day function to be uploaded when the actigraph is initialized and reduce the repetitive computational burden that would exist if a cosine table was used and the time of day function was calculated from the cosine table for each epoch.

The preferred embodiment, as shown in Figure 6, may also be realized by a stand-alone device or a component add-on to an actigraph 10. A suitable stand-alone device includes an input port (input means 20) to be physically connected to an input device, e.g., a keyboard, data entry device, or an actigraph through either a separate interface device or a cable/wire. Alternatively, this physical input connection could be replaced by a wireless communication system including telemetry, radio wave, infrared, PCS, digital, cellular, or light based system. The wireless communication system has an advantage in that it eliminates the need for a physical connection like cables/wires, plug-ins, etc. which is particularly convenient when monitoring a mobile subject. In accordance with an aspect of the invention, the actigraph may communicate via a wireless communication system to an external device that then is connected to the stand-alone device. The actigraph 10 preferably provides a sleep history that may include past, present and/or predicted/anticipated sleep patterns of an individual that may be analyzed by a sleep scorer 15 as illustrated in Figure 6. Alternatively, if the data from the actigraph 10 is not in the form of a sleep/wake history, then the actigraph data may be analyzed by software, for example from the manufacturer, to sleep score (sleep scorer 15) the actigraph data prior to input into the stand-alone apparatus.

The stand-alone device further includes an interpretation means 30 for analyzing and classifying the states present in the input data as usually either a sleep state or a wake state. Interpretation means 30 also selects or generates at least one of the following calculation functions responsive to the composition of the input data: 1) wake function, 2) sleep function, 3) delay function, and 4) sleep inertia function. Interpretation means 30 may be realized by an appropriately programmed integrated circuit (IC). One of ordinary skill in the art will realize that a

variety of devices may operate in concert with or be substituted for an IC like a discrete analog circuit, a hybrid analog/IC or other similar processing elements.

The stand-alone device further includes a determination means 40. Determination means 40 may be implemented by appropriately programming the  
5 IC of the interpretation means or it may be implemented through a separate programmed IC determining and calculating the cognitive performance capacity factoring in the sleep/wake history and the current state.

The interpretation means 30 and determination means 40 may be combined into one combined means or apparatus.

10 The calculated cognitive performance capacity may be stored in a first memory area 60 that stores modulation data including a modulating data series or curve preferably in the form of a time of day curve. The stand-alone device further includes a second memory area 50 that holds data for the creation of a data series or a curve representing cognitive performance capacity  $C$  over time  $t$ . The first  
15 memory 60 and the second memory 50 may be any memory known to those of ordinary skill in the art. The second memory 50 is preferably a first-in-first-out memory providing means for adding the value from the determination means 40 to the end of the data series or the curve. The first memory and the second memory may be combined as one memory unit.

20 The stand-alone device also includes, as a separate IC or in combination with one of the previously mentioned ICs, a modulation means 70 to modulate the first data series or curve (cognitive performance capacity) with the modulating data series or curve (time of day). The modulation preferably is performed by matching the timing sequence information relating to the data series or the curves based on  
25 length of time from initial input of data as preferably determined by the number of epochs and the initial starting time related to the first sleep/wake state.

The stand-alone device may also include a display to show a plotted modulated curve representing the modulation result over time or a numerical representation of a point on the modulated curve at a selected time from the  
30 modulation means 70 representing the predicted cognitive performance  $E$ . The numerical representation may take the form of a gauge similar to a fuel gauge in a motor vehicle. The stand-alone device, as an alternative or in addition to the display, may include a printer or communication port to communicate with an external device for printing and/or storage of the modulated curve or data series.

The stand-alone device instead of having dedicated hardware may provide the storage space and processing power to execute a software program and accompanying data files. In this case, the stand-alone device may be a desktop computer, a notebook computer, or similar computing device. The software  
5 program handles the receiving of the data representing sleep history from an outside source through a communication port, and then performs the necessary analysis and processing of the method described herein. The storage space may be a memory in the form of computer readable material storing at least the time of day curve and possibly the input data, which may also be resident in the random-  
10 access-memory (RAM) of the computer given its temporary use. The input data and the resulting produced data indicating various cognitive performance levels of an individual may also be saved to a more permanent memory or storage than RAM.

Another alternative embodiment provides for the interpretation means 30 to  
15 filter the sleep/wake data such that for the first  $k_d$  number of sleep epochs after a wake epoch are changed to wake epochs. In keeping with the invention, the filtering may be accomplished a variety of ways. The preferred way is to add a decision step prior to S3 in Figure 4 such that if  $D_{sw}$  is a sleep epoch and  $t - t_{LS} \leq k_d$ , then S3-S6a will be skipped and S7a will occur. The result is that the decision  
20 rule represented as  $d(t)$  in Equation 6 above would be eliminated, and S6b and S7d would be unnecessary in Figures 4(a)-(b) and 7(a)-(b).

The component add-on to an actigraph with sleep scoring capabilities may have similar components to the stand-alone device described above and shown in Figure 6. Preferably the component add-on is contained in one integrated chip to  
25 minimize the space needed to house it. However, the component add-on may include more than one electrical component, e.g., a memory chip and an IC. The component add-on may transmit the predicted cognitive performance to a remote device for further analysis.

Both the software and hardware are envisioned as being able to operate  
30 and function in real-time. For the purposes of this invention, real-time is understood to represent a continuous stream and analysis of cognitive performance level for each epoch of sleep/wake data entered. Thus, the software and hardware will both provide to an individual or some other entity the present cognitive performance level based on the data from the last entered epoch of



sleep/wake data entered into either the software or hardware. Most sleep scoring systems make the sleep/wake determination based on data from epochs on either side of the epoch being analyzed. Consequently, there is a delay in providing information to the user.

5       As one of ordinary skill in the art will appreciate from the following discussion, the described method is able to accept a continuous stream of data from either individual epochs or groups of epochs. If blocks of time are entered, then after initial transitions the first few epochs are governed by the appropriate transition function with the appropriate time of solid sleep or wakefulness being  
10       used in the non-transition functions.

      As a feature of the invention, the sleep/wake data may comprise the time at which a state change occurs from sleep to wake or wake to sleep. The sleep/wake data may also comprise the duration of the individual's wake state and the duration of the individual's sleep state. In order to generate the predicted cognitive  
15       performance curve, the sleep/wake data may be extrapolated and/or expanded into a series of individual epochs. As discussed above an epoch represents a predetermined length of time. Thus the sleep/wake data may be presented in conventional units of time or may be presented in epochs. For example, if the sleep/wake data was sleep for 10 epochs and wake for 3 epochs, in generating the  
20       cognitive performance capacities, epochs 1 through 10 may represent the sleep state and epochs 11 through 13 may represent the wake state.

      In accordance with an aspect of the invention, the predicted cognitive performance  $E$  at a particular time  $q$  may be determined using either the predicted cognitive performance  $E$  or the cognitive performance capacity  $C$  at time  $r$  as a  
25       base point where  $r$  can be before or after time  $q$ . From the base point determining the cognitive performance capacities for the time points between times  $q$  and  $r$  where there is a change in state.

      As shown in Figures 7(a)-(b), the steps are substantially the same as the preferred embodiment with changes made to the wake and sleep functions,  
30       consequently the definition of the variables is the same as the preferred embodiment except as noted. The equations described below and the steps shown in Figures 7(a)-(b) are for the situation when the initial cognitive value is prior in time to the desired predicted cognitive value. Each element of sleep/wake data is classified as either sleep or wake.

If the sleep/wake data represents the wake state, then a selection is made between two functions as to which is applicable based on the following decision rule:

IF  $\Delta t \leq k_i$   
 5 THEN  $C_t = i(t)$   
 ELSE  $C_t = w_m(t)$

Equation 10

where  $\Delta t$  represents the amount of time in the current state, i.e.,  $t - t_{LS}$ . The sleep inertia function  $i(t)$  is used only if the last data entry is the wake state for a period of time is less than or equal to  $k_i$ . Thus the same sleep inertia function  $i(t)$  as used in  
 10 the preferred embodiment is also used in this alternative embodiment. The modified wake function  $w_m(t)$  takes into account that the sleep inertia function  $i(t)$  provides a delay of  $k_i$  when a curve is formulated, such that after an individual recovers from the initial suppression of cognitive performance capacity the individual returns to the level of cognitive performance capacity of the last epoch  
 15 the individual was asleep prior to waking. Accounting for this delay provides the following:

$$w_m(t) = C_{t-1} - k_w(\Delta t - k_i)$$

Equation 11

If the sleep/wake data represents the sleep state, then a selection is made between two functions as to which is applicable based on the following decision  
 20 rule:

IF  $\Delta t \leq k_d$   
 THEN  $C_t = d(t)$   
 ELSE  $C_t = s_m(t)$

Equation 12

The delay function  $d(t)$  is used only if the last data entry is the sleep state for a  
 25 period of time is less than or equal to  $k_d$ . Thus the same delay function  $d(t)$  as used in the preferred embodiment is also used in this alternative embodiment. The modified sleep function  $s_m(t)$  takes into account the delay function for a period of time equal to  $k_d$ . Accounting for the delay function  $d(t)$  provides the following:

$$s_m(t) = ((C_{t-1} - (k_w * k_d)) + (100 - (100 - C_{t-1}) (1 - 1/k_s)^{\Delta t - k_d}))$$

Equation 13

30 where the first part of the equation represents the delay function  $d(t)$  and the second part represents the recovery of cognitive performance capacity  $C$ .

A summation of the time components of the sleep/wake data is performed as each piece of sleep/wake data is handled with respect to the calculation of the cognitive performance capacity or prior to modulation of the final cognitive

If the sleep/wake data represents the wake state, then a selection is made between two functions as to which is applicable based on the following decision rule:

$$\begin{aligned}
 & \text{IF } \Delta t \leq k_i \\
 5 \quad & \text{THEN } C_t = i(t) \\
 & \text{ELSE } C_t = w_m(t)
 \end{aligned}
 \tag{Equation 10}$$

where  $\Delta t$  represents the amount of time in the current state, i.e.,  $t - t_{LS}$ . The sleep inertia function  $i(t)$  is used only if the last data entry is the wake state for a period of time is less than or equal to  $k_i$ . Thus the same sleep inertia function  $i(t)$  as used in the preferred embodiment is also used in this alternative embodiment. The modified wake function  $w_m(t)$  takes into account that the sleep inertia function  $i(t)$  provides a delay of  $k_i$  when a curve is formulated, such that after an individual recovers from the initial suppression of cognitive performance capacity the individual returns to the level of cognitive performance capacity of the last epoch the individual was asleep prior to waking. Accounting for this delay provides the following:

$$w_m(t) = C_{t-1} - k_w(\Delta t - k_i) \tag{Equation 11}$$

If the sleep/wake data represents the sleep state, then a selection is made between two functions as to which is applicable based on the following decision rule:

$$\begin{aligned}
 & \text{IF } \Delta t \leq k_d \\
 20 \quad & \text{THEN } C_t = d(t) \\
 & \text{ELSE } C_t = s_m(t)
 \end{aligned}
 \tag{Equation 12}$$

The delay function  $d(t)$  is used only if the last data entry is the sleep state for a period of time is less than or equal to  $k_d$ . Thus the same delay function  $d(t)$  as used in the preferred embodiment is also used in this alternative embodiment. The modified sleep function  $s_m(t)$  takes into account the delay function for a period of time equal to  $k_d$ . Accounting for the delay function  $d(t)$  provides the following:

$$s_m(t) = ((C_{t-1} - (k_w * k_d)) + (100 - (100 - C_{t-1}) (1 - 1/k_s)^{\Delta t - k_d})) \tag{Equation 13}$$

where the first part of the equation represents the delay function  $d(t)$  and the second part represents the recovery of cognitive performance capacity  $C$ .

A summation of the time components of the sleep/wake data is performed as each piece of sleep/wake data is handled with respect to the calculation of the cognitive performance capacity or prior to modulation of the final cognitive

performance capacity with the time of day function  $m(t)$ . The latter is shown in Figures 7(a)-(b). After the new cognitive performance capacity  $C_t$  is calculated, the method repeats to handle the next piece of sleep/wake data if the present piece is not the last piece. After the last piece the predicted cognitive performance  $E$  is  
5 calculated based on Equation 1 above and as detailed in the preferred embodiment.

It should be noted again that this method includes the processes and calculations based on Equations 1 through 8 expressed in their general form. Embodiments shall apply functions relating the variables involved according to  
10 empirical knowledge, resulting in specific expressions of those equations, as illustrated in the text and Figures 1-7(b) above (but not confined to these), which may be changed or refined according to the state of empirical knowledge.

#### **Applications of the Method**

For the discussions that follow about Figures 8-17, the predicted cognitive  
15 performance  $E$  (i.e., the combination of cognitive performance capacity  $C$  and time of day function  $M$ ) is plotted as a continuous line across multiple days. The darker sections of the line indicate sleep periods and the lighter sections of the line indicate wake periods. Predicted cognitive performance  $E$  is illustrated on a scale of 0 to 120%. Using the preferred embodiments, predicted cognitive performance  
20  $E$  can theoretically reach 120%, but only when cognitive performance capacity  $C$  is 100% (i.e., 20 minutes after awakening from a sleep period in which cognitive performance capacity  $C$  was fully restored) and simultaneously the time of day function  $M$  is at its acrophase. Although possible, in practice this situation is unlikely. For purposes of illustration and description, "acceptable" predicted  
25 cognitive performance  $E$  is set at 85%. This value represents the approximate percent decrement in predicted cognitive performance  $E$  after 16 hours of continuous wakefulness when preceded by 8 hours of nighttime sleep (i.e., a typical sleep/wake schedule). Temporal resolution along the abscissa of the illustrations is one hour per vertical tick mark. Major vertical gridlines correspond to  
30 0:00 AM (midnight) local time.

##### **(1) Impact of Idealized Sleep on Predicted Cognitive Performance $E$**

In its simplest application, the method according to the invention may be used to predict the impact of various idealized (i.e., unfragmented) amounts of nightly sleep on predicted cognitive performance  $E$ . For this exercise, the

individual obtains eight or four hours, respectively, of unfragmented sleep per night for the first night. Next, the individual obtains 8 (Figure 8) or 4 (Figure 9) hours of unfragmented sleep per night. Then the individual again obtains eight hours of unfragmented sleep.

5       As illustrated in Figure 8, the method predicts that with eight hours of sleep per night, acceptable predicted cognitive performance E is maintained across waking hours, dropping only slightly below 85% for the 40 minutes preceding sleep onset each day. As illustrated in Figure 9, for sleep amounts of four hours per night, the method predicts that after the first night of restricted sleep, predicted  
10       cognitive performance E falls below acceptable levels for the entire waking period. Furthermore, because of the restricted sleep period, predicted cognitive performance E is not completely restored each night, despite the higher rate of accumulation of cognitive performance E during sleep.

Under conditions of total sleep deprivation illustrated in Figure 10  
15       commencing on night 2, the method predicts that predicted cognitive performance E falls to near-zero levels on Day 4 after 88 hours awake. The predicted rate of cognitive performance E accumulation during recovery sleep is shown in Figure 11. This steep rate of accumulation on the first night (Day 2) of recovery sleep is a result of the near-complete depletion of predicted cognitive performance E.  
20       However, in spite of this steep rate, predicted cognitive performance E accumulation is not complete (above the arbitrary 85% "acceptable" level) until after the second night (Day 3) of recovery sleep. Likewise, the rate of accumulation on the second night of recovery sleep is slightly lower than the rate on the first night of recovery sleep, due to the lower level of preceding sleep debt  
25       (and correspondingly higher level of predicted cognitive performance E) at sleep onset.

The impact of a near-complete depletion of predicted cognitive performance E on the rate of predicted cognitive performance E accumulation during sleep is illustrated in Figure 12. This figure shows predicted cognitive performance E  
30       across 3.5 days of sleep deprivation in which a daily 30-minute sleep period occurs on Days 2-4. The rate of accumulation during the 30-minute sleep period is nearly vertical. Although predicted cognitive performance E declines both within and across each day, it is substantially offset by the 30-minute sleep period.

(2) Impact of Sleep Fragmentation on Predicted Cognitive Performance E

Another practical application uses the method to predict the cognitive performance in an individual with fragmented sleep, either due to a sleep disorder such as sleep apnea or due to environmental disturbances such as airplane or train noises. Figure 13 illustrates predicted cognitive performance E across three nights of fragmented sleep, during which arousals (brief awakenings) occur 10 times per hour. Daytime predicted cognitive performance E following nights on Days 2-4 (when sleep is disturbed) is severely impaired, and continues to decline across each day. Predicted cognitive performance E will not be completely restored even after one night of eight hours of undisturbed sleep.

(3) Predicted Cognitive Performance E across Two Night Shifts

Another practical application uses the method to predict the cognitive performance E of an individual across two nights of shift work. Figure 14 illustrates predicted cognitive performance E across two nights during which the individual works from 11:00 PM to 7:00 AM and sleeps from 8:00 AM to 4:00 PM. Prior to the first shift, the individual sleeps his usual day-shift hours, i.e., from midnight to 8:00 AM. He stays up all day and starts his first shift at 11:30 PM that night. The shaded area shows that he is working when predicted cognitive performance E is poorest for that 24-hour period. This is due to the combined influences of sleep deprivation and time of day. The person then sleeps from 8:00 AM to 4:00 PM -- for the purposes of this exercise, it is assumed that the person actually slept the entire eight hours of this latter sleep period. Substantial recovery of predicted cognitive performance E is obtained during this sleep period. The second shift begins with predicted cognitive performance E at near-optimal levels. However, due to time of day effects, the bulk of the second shift occurs when the method predicts poorest cognitive performance E.

(4) Retrospective Analysis of Predicted Cognitive Performance E

In another application, the method is used to retrospectively predict cognitive performance E in a commercial motor vehicle operator involved in a driving collision/traffic accident. The driver's approximate sleep and wake times (taken from personal history) for several days prior to the collision serve as input. Predicted cognitive performance E based on this approximate sleep/wake data is depicted in Figure 15. The driver's trip began the morning of Day 1. Due to an early start time, the driver began the trip partially sleep deprived. Consequently,

his predicted cognitive performance E at the start of the trip was only slightly above the arbitrarily defined acceptable level of 85%. During the first day (Day 1) on the road, the driver worked for 16 hours – predicted cognitive performance E during the latter portion of this period fell below acceptable levels. The driver stopped for his first sleep at 4:00 AM on Day 2. Sleep on Days 2-4 was of insufficient duration to restore predicted cognitive performance E. On Day 5, the driver obtained little sleep (30 minutes). The collision occurred later that day as the driver's predicted cognitive performance E began the rapid evening decline. At the time of the collision, the driver's predicted cognitive performance E was approximately 50% of optimal levels. Finally, given the extreme sleep deprivation, the predicted cognitive performance recovery slope during sleep on Day 6 (following the collision) was steep, restoring predicted cognitive performance E to near the arbitrary 85% "acceptable" level.

(5) Predicted Cognitive Performance E Based on Current Sleep/Wake Schedule and Modification of Future Sleep/Wake to Optimize Predicted Cognitive Performance E

In this application, the method is used first to predict an individual's level of cognitive performance E across some interval based on that individual's current work and sleep/wake schedule. Next, the method is used to re-schedule sleep and wakefulness in order to optimize predicted cognitive performance E over the same interval.

In this example, first we model a driver's predicted cognitive performance E based on his current sleep/wake schedule. The driver's current sleep/wake schedule is generated around the maximum duty hours allowed under the Federal Highway Administration's (FHWA) hours-of-service regulations. These regulations allow the driver to obtain a maximum 15 hours on-duty (maximum 10 hours driving plus five hours on-duty but not driving) followed by a minimum eight hours off-duty. The driver may continue this on/off-duty cycling until 60 hours on-duty has been accumulated—at which point the driver must take time off until seven days has elapsed since he commenced duty. Figure 16 illustrates the driver's sleep/wake schedule and predicted cognitive performance E under these restrictions. The driver sleeps six of his eight hours off-duty. The schedule results in a 23-hour "day" which means that the driver initiates sleep one hour earlier each evening. Because partial sleep deprivation occurs and sleep is timed earlier each day, the

invention predicts that by the second duty day the driver will spend a substantial portion of on-duty time with predicted cognitive performance E below the arbitrarily defined 85% cut-off. When the driver reaches the maximum 60 hours on-duty, he must then take several days off even though the invention predicts that cognitive performance E is restored after only one night of 10 hours sleep.

Figure 17 illustrates an alternative work schedule also allowed under current FHWA regulations. This work schedule is based on a schedule of 12 hours on-duty and 12 hours off-duty. It is assumed that the driver sleeps eight of his 12 hours off-duty. Because no sleep deprivation or shifting of sleep timing occurs, the invention predicts that the driver maintains cognitive performance E at or above 85% within and across the duty days.

## VI. INDUSTRIAL APPLICABILITY

Although described above in connection with a variety of specific activities, this invention has many other applications. The method for predicting cognitive performance will provide critical information for managing both individual and group productivity. For example, in military operational planning, this method will enable commanders to determine precisely, based on prior sleep history, each soldier's current and predicted level of cognitive performance. Commanders can also input a likely sleep/wake schedule and thereby predict a soldier's cognitive performance throughout an impending mission. Throughout conduct of the mission itself, the latter cognitive performance predictions (originally based on likely sleep/wake schedule) can be updated based on actual sleep acquired. The ability to project future cognitive performance will allow commanders to optimize troop performance during continuous operations by, for example, planning sleep/wake schedules around the mission to optimize cognitive performance, selecting those troops or combinations of troops whose predicted cognitive performance will be maximal at a critical time, etc. This method will assist in maximizing productivity at both the individual and unit level.

This invention may be employed in a variety of commercial applications covering many occupational areas for purposes of optimizing output (productivity). The invention provides managers with the capability to plan operations and regulate work hours to a standard based on objective cognitive performance predictions. This is in contrast to the frequently used method of regulating work hours by time off-duty (a relatively poor predictor of sleep/wake patterns and



consequently a poor predictor of cognitive performance) or by generating alertness/sleepiness predictions (which, as noted above, do not always correspond to cognitive performance). The invention can be "exercised" in hypothetical sleep/wake scenarios to provide an estimate of cognitive performance under such scenarios. To the extent that optimizing cognitive performance is of interest to the general public, there is a possibility for use in a variety of applications.

The method may also be used to gauge and evaluate the cognitive performance effects of any biomedical, psychological, or other (e.g., sleep hygiene, light therapy, etc.) treatments or interventions shown to improve sleep. Examples of these include but are not limited to patients with overt sleep disorders, circadian rhythm disorders, other medical conditions impacting sleep quality and/or duration, poor sleep hygiene, jet lag, or any other sleep/wake problem. Currently, the efficacy of treatments for improving sleep is determined by comparing baseline polysomnographic measures of nighttime sleep and some measure of daytime alertness (e.g., the MSLT, the Maintenance of Wakefulness Test (MWT), the Stanford Sleepiness Scale or the Karolinska Sleepiness Scale) with the same measures obtained after treatment. Both treatment efficacy and the likely impact on performance during waking periods are inferred from the results on the daytime alertness tests. For example, the Federal Aviation Administration currently requires any commercial pilots diagnosed with sleep apnea to undergo treatment. Such treatment is followed by daytime alertness testing on a modified version of the MWT. During the MWT, pilots are put in a comfortable chair in a darkened room and instructed to try to remain awake for extended periods. If the pilots are able to avoid overt sleep under these sleep-conducive conditions then they are deemed fit for duty. The inference is that the minimal ability to maintain wakefulness at a discrete point in time translates into the ability to operate an aircraft safely (i.e., it is inferred that alertness is equivalent to cognitive performance). However, sleep deprivation can affect cognitive performance even when it does not result in overt sleep, particularly during an alertness test when for various reasons the individual may be highly motivated to stay awake.

In contrast, the current method allows cognitive performance to be estimated directly from measured sleep parameters considered in conjunction with the time of day. The advantages of this method over current methods for evaluating treatment efficacy are: (1) the motivations and motivation levels of the

patients being tested cannot affect results (cognitive performance determinations); and (2) the method allows numerical specification and prediction of cognitive performance across all projected waking hours rather than indicating alertness at a discrete, specified point in time. Thus, the method provides a continuous scale for gauging cognitive performance across time rather than providing only a minimal "fitness for duty" determination based on the patient's ability to maintain EEG-defined wakefulness at a specific time.

The method may also be used clinically as an adjunct for diagnosing sleep disorders such as narcolepsy and idiopathic CNS hypersomnolence. Equally important, it may also be used to differentiate among sleep disorders. The latter is critical to the course of treatment, and consequent treatment efficacy depends on a valid and reliable diagnosis. For example, sleep apnea and periodic limb movements during sleep are characterized by nighttime sleep disruption (i.e., partial sleep deprivation) accompanied by daytime cognitive performance deficits. In contrast, narcolepsy and idiopathic hypersomnolence tend to be characterized by apparently normal nighttime sleep, but accompanied by daytime cognitive performance deficits. Based on the apparently normal nighttime sleep in the latter two groups, the invention would predict relatively normal cognitive performance. Thus, a discrepancy between predicted cognitive performance (based on the current invention) and observed or measured cognitive performance could be used to distinguish one sleep disorder from another. For example, narcolepsy, idiopathic hypersomnolence, or other CNS-related causes of daytime cognitive performance deficits (where no sleep deficit is apparent) could be distinguished from sleep apnea, periodic limb movements, or other causes of daytime cognitive deficits (where impaired sleep is evident).

Those skilled in the art will appreciate that various adaptations and modifications of the above-described preferred embodiments can be configured without departing from the scope and spirit of the invention. Therefore, it is to be understood that, within the scope of the appended claims, the invention may be practiced other than as specifically described herein.

**IN THE CLAIMS:**

We claim

1. A method comprising:  
collecting activity information of an individual with an actigraph,  
providing a data series representing wake states and sleep states of the individual based on an analysis of the activity information using a sleep scoring  
5 system,  
selecting a function based on the type of data in the data series,  
calculating a cognitive performance capacity based on the selected function,  
modulating the cognitive performance capacity with a time of day value, and  
outputting the modulated value as the cognitive performance level.
2. The method according to claim 1, further comprising:  
storing the modulated values as coordinates representing time and  
amplitude,  
repeating the calculating, modulating and outputting steps of claim 1,  
5 plotting a curve from the stored modulated values, and  
outputting the curve representing cognitive performance level over a period  
of time.
3. The method according to claim 2, further comprising extrapolating  
from the curve a predictive curve based on anticipated wake states and anticipated  
sleep states.
4. The method according to claim 1, wherein said outputting step  
includes outputting the modulated value to a display.
5. The method according to claim 1, wherein said outputting step  
includes outputting the modulated value to a data file.
6. The method according to claim 1, wherein said outputting step  
includes outputting the modulated value to a printing device.
7. The method according to claim 1, further comprising formulating the  
time of day values to represent a curve having a period of 24 hours such that the  
curve includes a first sinusoidal curve having a 24-hour period and a second  
sinusoidal curve having a 12-hour period.
8. The method according to claim 1, wherein the time of day values  
represent a curve having a period of 24 hours such that the curve includes a first

sinusoidal curve having a 24-hour period and a second sinusoidal curve having a 12-hour period.

9. The method according to claim 1, wherein the selecting step includes selecting from a group consisting of a wake function, a sleep function, a delay function, and a sleep inertia function.

10. The method according to claim 1, wherein the selecting step includes determining the present state for the data series as either the wake state or the sleep state,

calculating a length of time in the present state, and

5 selecting the function based on one of:

the length of time in the present state and the present state, and

the length of time in a recent transition from wake to sleep or sleep to wake.

11. The method according to claim 10, wherein the selecting step includes selecting the function from a group consisting of a wake function, a sleep function, a delay function, and a sleep inertia function.

12. The method according to claim 1, wherein the calculating step calculates a cognitive performance level as a percentage value such that 100% is a maximum cognitive performance capacity.

13. The method according to claim 1, wherein said method is performed in real-time.

14. A method comprising:

collecting activity information of an individual with an actigraph,

providing a data series representing wake states and sleep states of the individual based on an analysis of the activity information using a sleep scoring  
5 system,

selecting a function based on the type of data in the data series,

calculating a cognitive performance capacity based on the selected function,

approximating a first curve of calculated cognitive performance capacities,

storing the first curve,

10 modulating the first curve with a second curve representing time of day rhythms, and

outputting the modulated first curve.

15. The method according to claim 14, wherein said outputting step includes outputting a value of a point on the modulated first curve to a display.

16. The method according to claim 14, wherein said outputting step includes outputting a value of a point on the modulated first curve to a data file.

17. The method according to claim 14, wherein said outputting step includes outputting a value of a point on the modulated first curve to a printing device.

18. The method according to claim 14, further comprising extrapolating from the modulated first curve a predictive curve based on anticipated wake states and sleep states.

19. The method according to claim 14, further comprising formulating the second curve having a period of 24 hours such that the curve includes a first sinusoidal curve having a 24-hour period and a second sinusoidal curve having a 12-hour period.

20. The method according to claim 14, wherein the second curve having a period of 24 hours such that the curve includes a first sinusoidal curve having a 24-hour period and a second sinusoidal curve having a 12-hour period.

21. The method according to claim 14, wherein the selecting step includes selecting from a group consisting of a wake function, a sleep function, a delay function, and a sleep inertia function.

22. The method according to claim 14, wherein the selecting step includes:

determining the present state for the data series as either the wake state or the sleep state,

5 calculating a length of time in the present state, and

selecting the function based on one of

the length of time in the present state and the present state, and

the length of time in a recent transition from wake to sleep or sleep to wake.

23. The method according to claim 22, wherein the selecting step includes selecting the function from a group consisting of a wake function, a sleep function, a delay function, and a sleep inertia function.

24. The method according to claim 14, wherein the calculating step includes calculating a cognitive performance level as a percentage value such that 100% is a maximum cognitive performance capacity.

25. The method according to claim 14, wherein said method operates in real-time.

26. A system comprising:

an actigraph,

scoring means for providing a sleep score based on data collected by said actigraph,

5 input means for receiving data from said scoring means,

interpretation means for analyzing the received data and selecting a calculation function based on the composition of the received data,

determination means for calculating a value using the calculation function,

means for adding the value from said determination means to a curve

10 including prior values, and

modulating means for modulating the curve with a modulating curve to produce an output curve.

27. The system of claim 26, further comprising a display means for displaying the last value of the output curve.

28. The system of claim 26, wherein said interpretation means determines a time representing a length of time of a current state of the data.

29. The system of claim 26, wherein said interpretation means determines a time representing a transition from one state to another.

30. The system of claim 26, wherein the modulating curve represents variations over a 24-hour period.

31. The system of claim 26, wherein said modulating means matches the curve to the modulating curve based on time sequencing.

32. The system of claim 26, wherein

said actigraph includes

said scoring means, and

communication means for communicating with said input means, said

5 communication means connected to said scoring means; and

said input means receiving data from said scoring means through said communication means.

33. The system of claim 26, wherein said determination means selects the function from a group consisting of a wake function, a sleep function, a delay function, and a sleep inertia function.

34. The system of claim 26, further comprising storage means for storing the output curve from said modulating means.

35. The system of claim 34, wherein said storage means stores the output curve as a series of data.

36. The system of claim 26, further comprising a housing, wherein said actigraph, scoring means, input means, interpretation means, determination means, adding means, and modulating means are located within said housing.

37. The system of claim 36, further comprising  
means for collecting data, and  
means for communicating either the value from said determination means or the output curve to said collecting means, said communicating means attached to  
5 said housing.

38. The system of claim 36, wherein said collecting means includes analysis means for analyzing data collected from multiple actigraphs.

39. The system of claim 26, wherein said adding means is memory.

40. A system comprising:  
an actigraph,  
a scorer connected to said actigraph,  
an input connected to said scorer,  
5 a data analyzer connected to said input to select a calculation function responsive to data from said scorer,  
a calculator connected to said data analyzer to calculate a cognitive performance capacity using the calculation function,  
a first memory that stores modulation data, and  
10 a modulator connected to said first memory and said calculator to modulate the cognitive performance capacity with the modulation data to generate a predicted cognitive performance.

41. The apparatus of claim 40, further comprising a display connected to said modulator.

42. The apparatus of claim 40, further comprising an output connected to said modulator.

43. The apparatus of claim 40, further comprising a second memory connected to said modulator containing data.

44. The apparatus of claim 43, wherein said first memory and said second memory are integrally formed as one memory.

45. The apparatus of claim 43, wherein the modulation data represents time of day variations over a 24-hour period.

46. The system of claim 43, wherein said modulator matches the data in said second memory to the modulation data based on time sequencing.

47. The apparatus of claim 43, wherein said second memory is a first-in-first-out memory.

48. The apparatus of claim 40, wherein said input is a telemetric receiver.

49. The apparatus of claim 40, wherein said input is a keyboard.

50. The apparatus of claim 40, further comprising a first-in-first-out memory connected to said modulator that stores modulated data.

51. The system of claim 40, further comprising a housing, wherein said actigraph, scorer, said input, said data analyzer, said calculator, said first memory, and said modulator are within said housing.

52. The system of claim 51, further comprising

a central data unit, and

a transmitter connected with either said calculator or said modulator, said transmitter attached to said housing;

5 wherein said transmitter communicates with said central data unit.

53. The system of claim 52, wherein said central data unit stores and analyzes data collected from multiple actigraphs.



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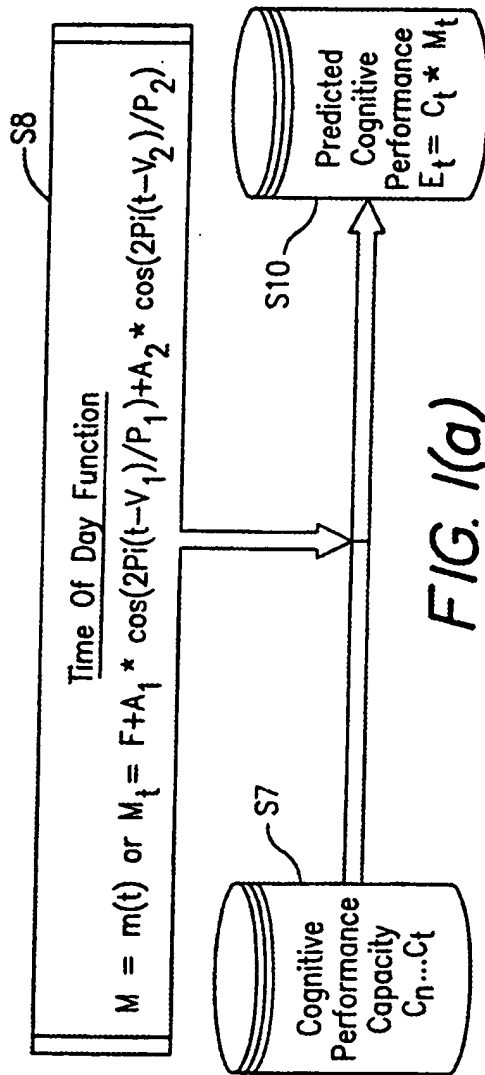


FIG. 1(a)

FIG. 1(b)

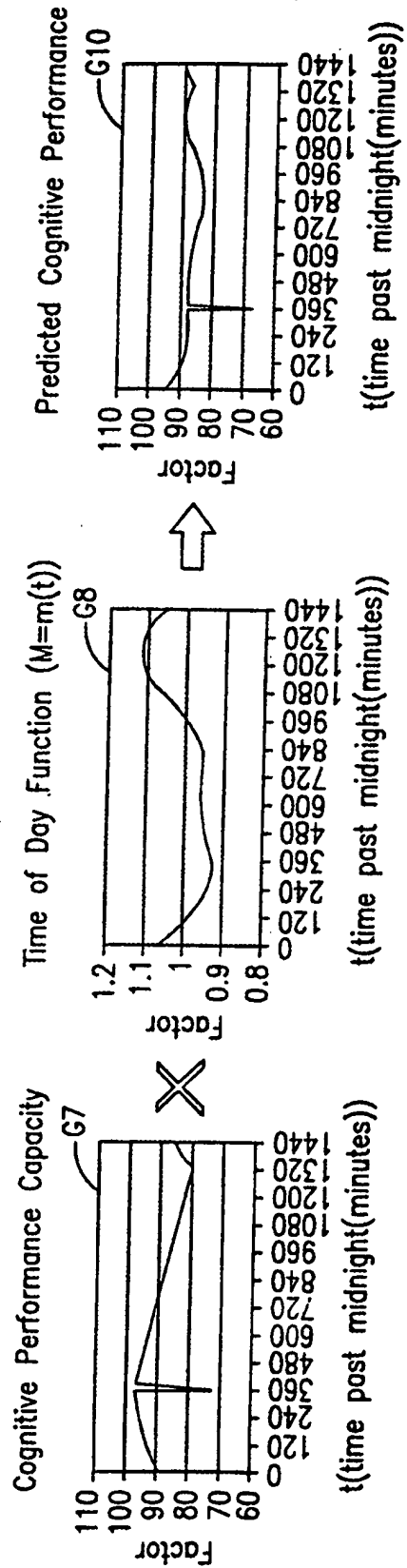


FIG. 2

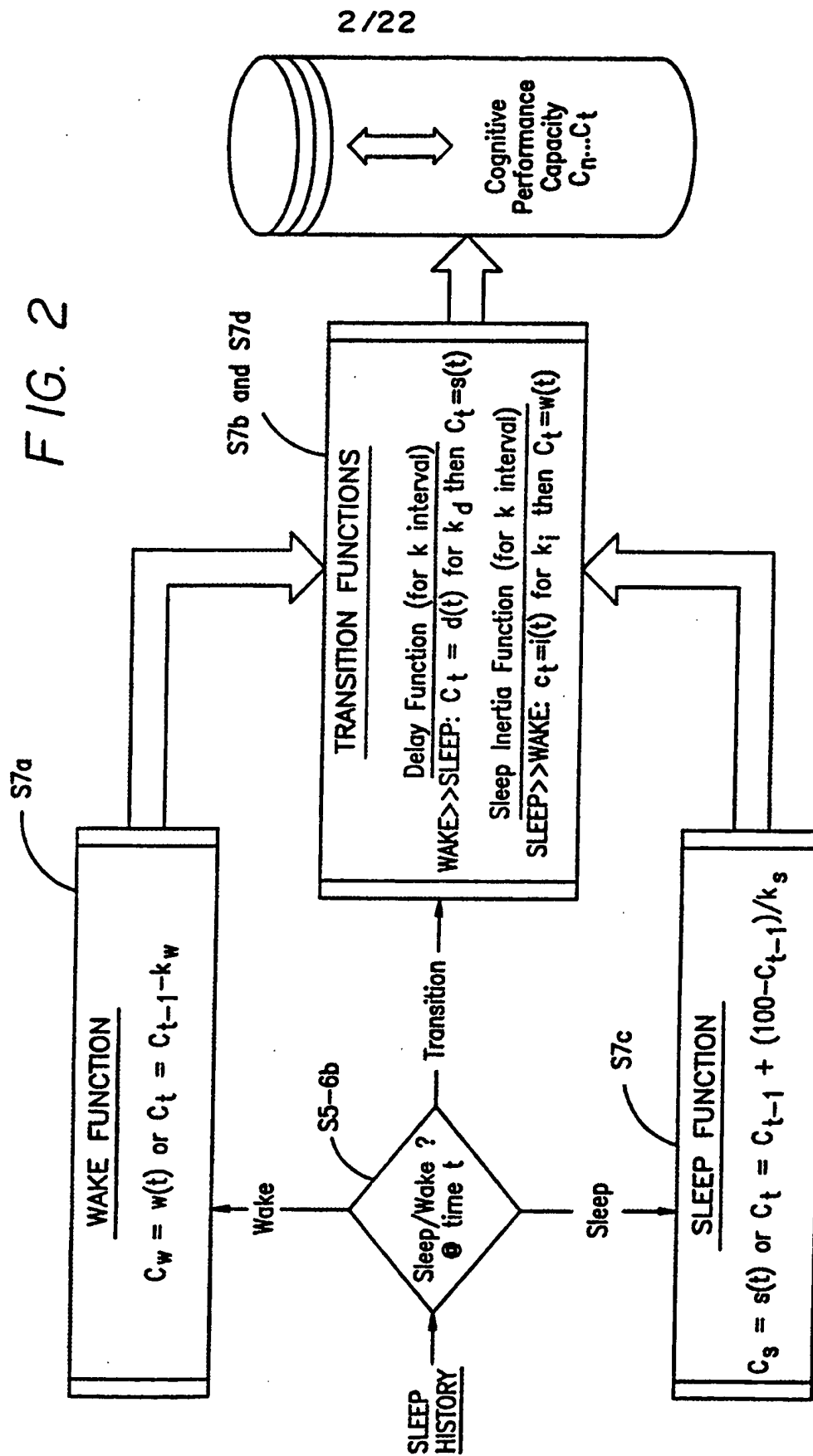


FIG. 3(a)

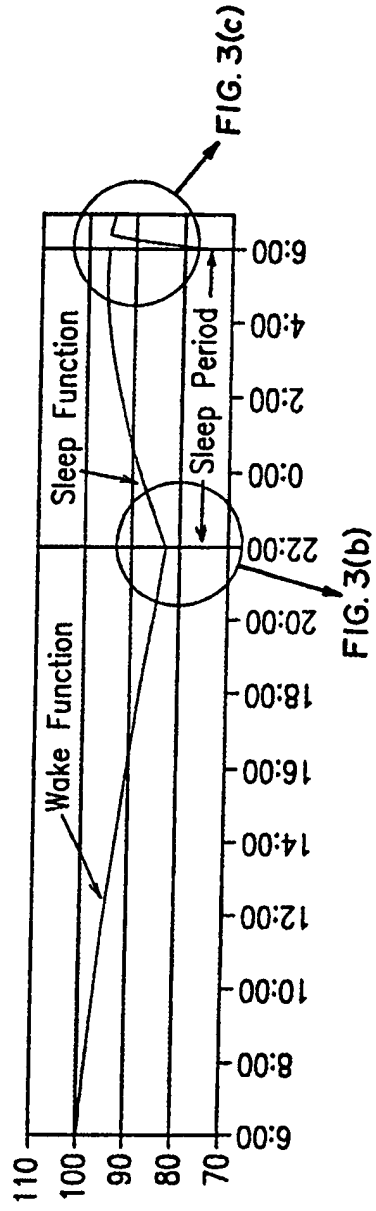


FIG. 3(b)

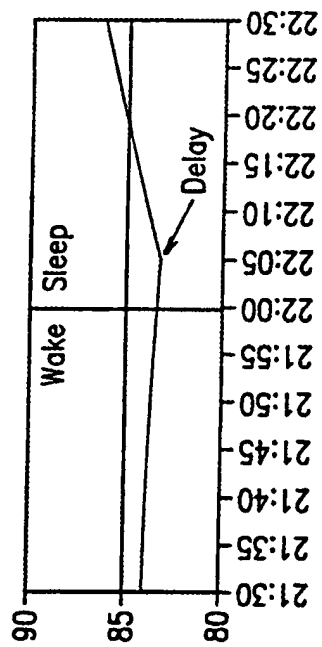
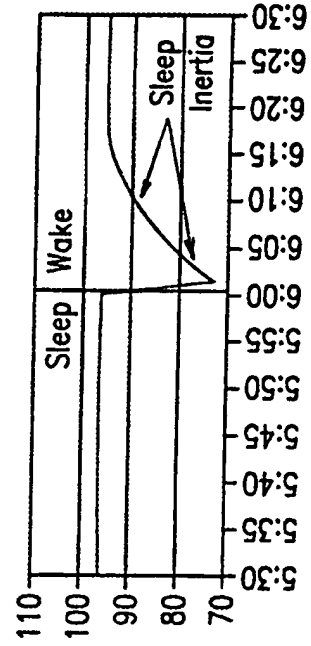
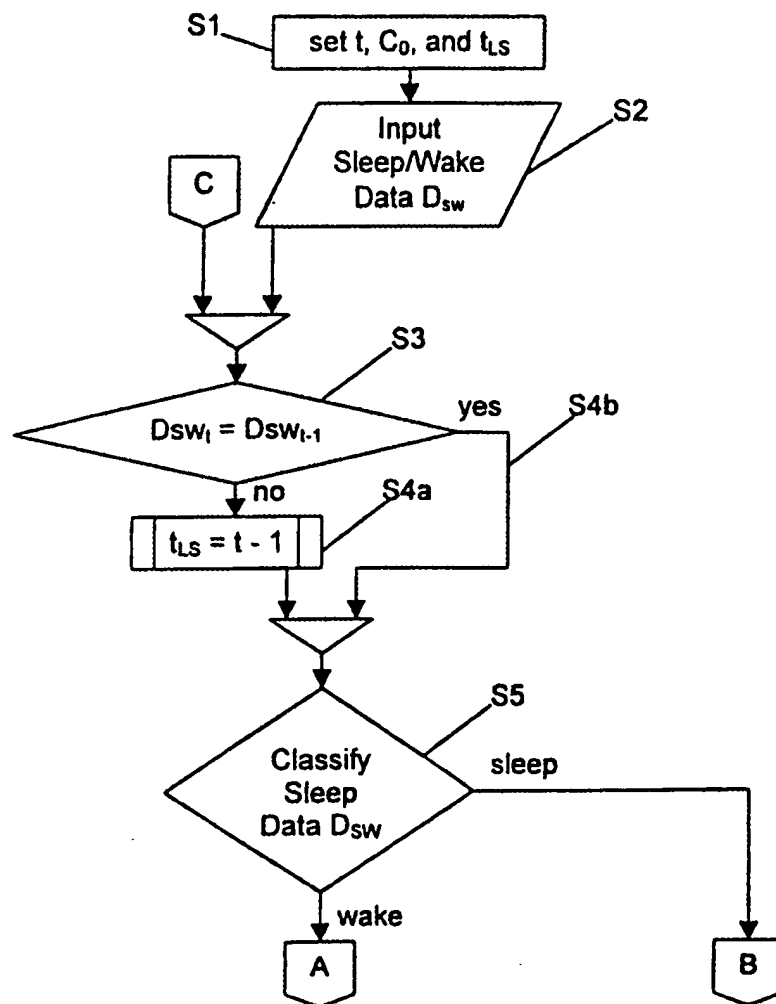


FIG. 3(c)



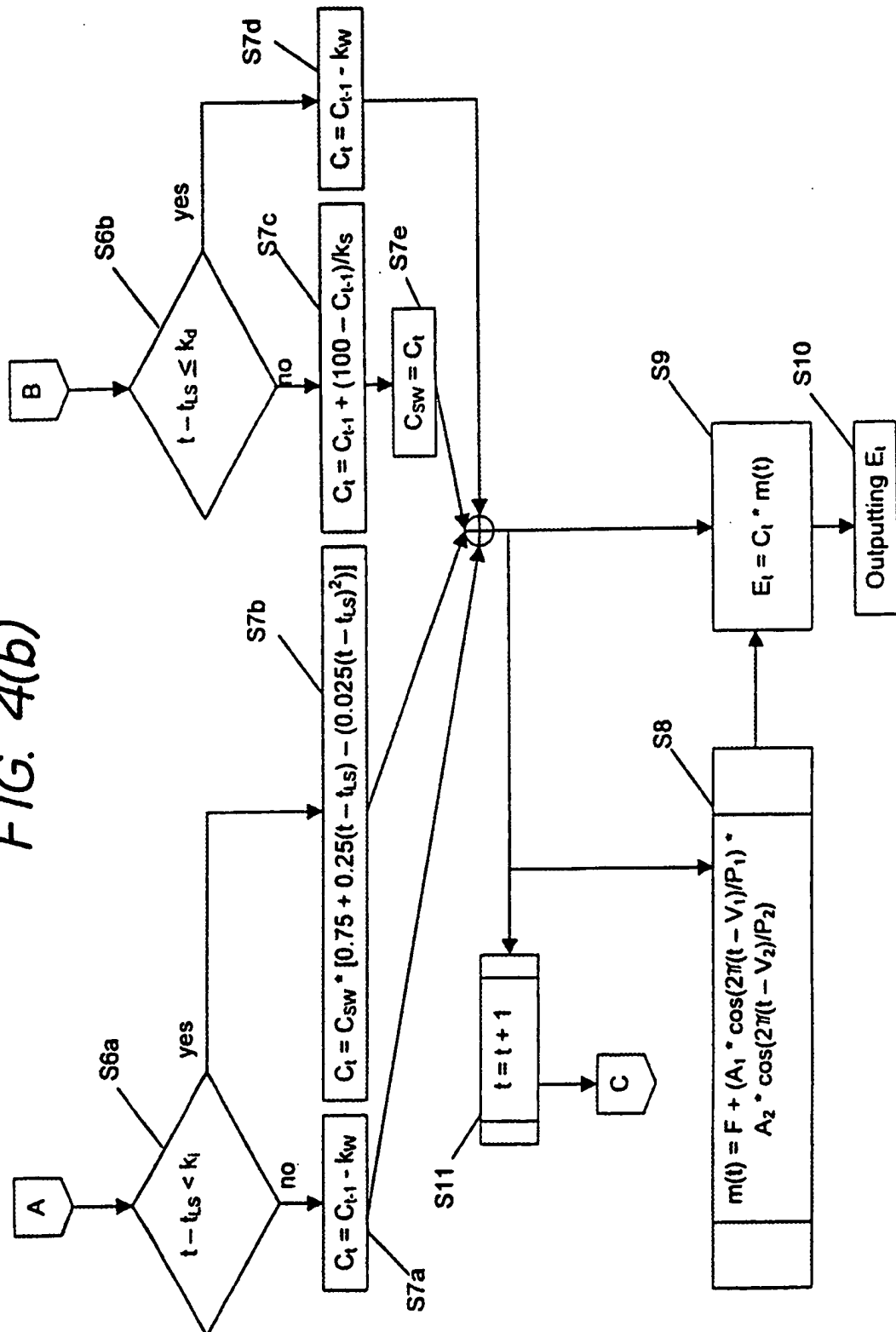
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FIG. 4(a)



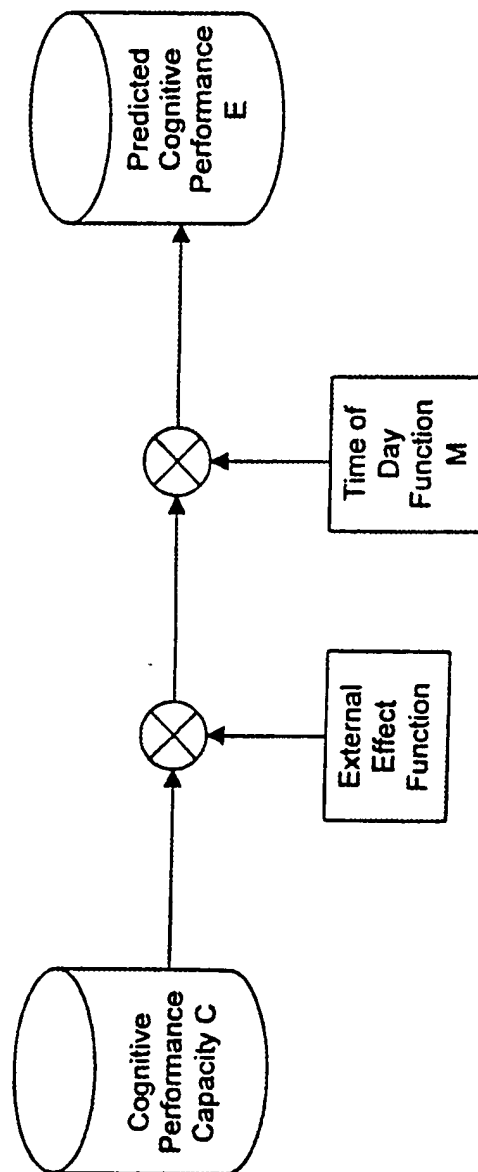
5/22

FIG. 4(b)



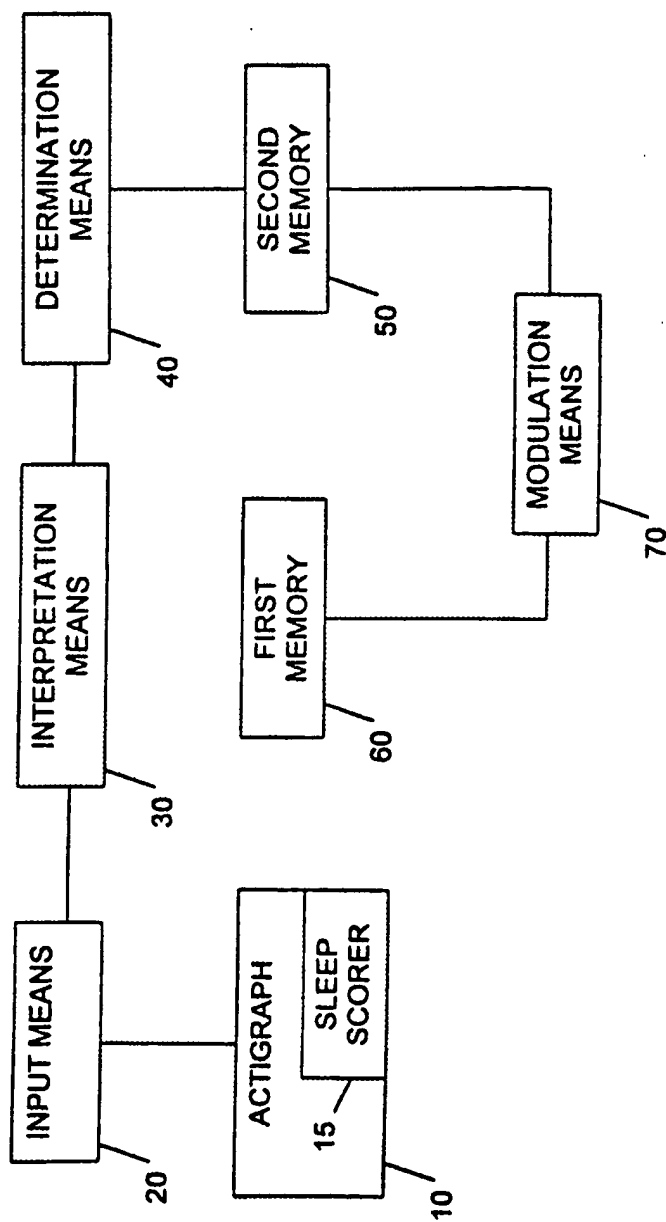
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FIG. 5



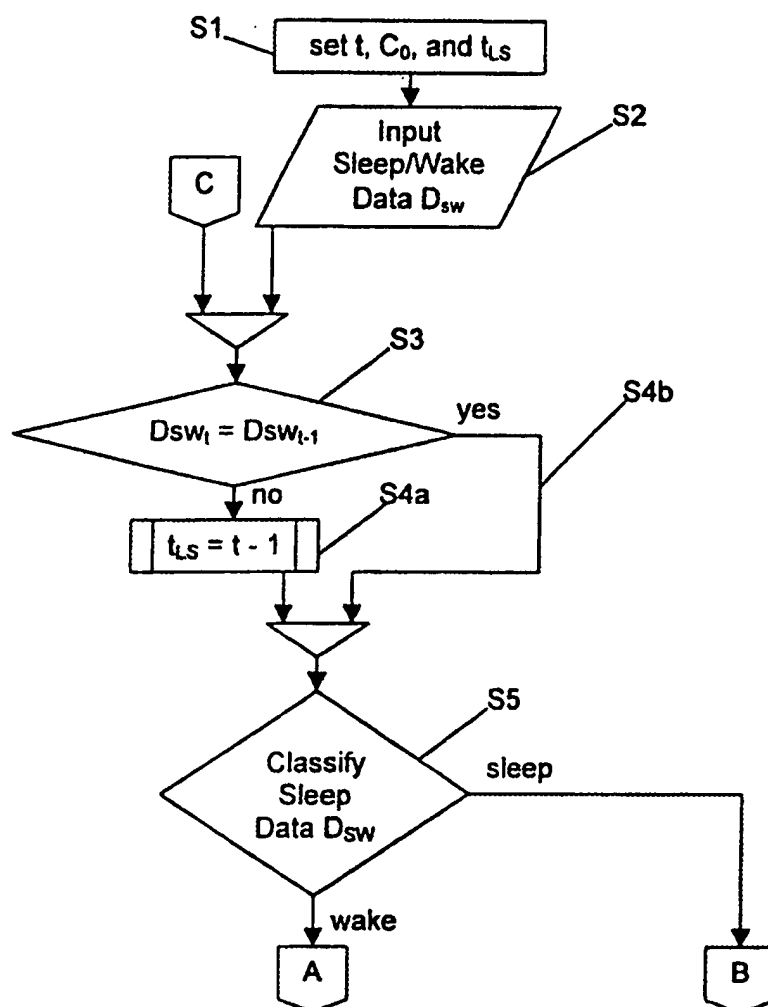
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FIG. 6



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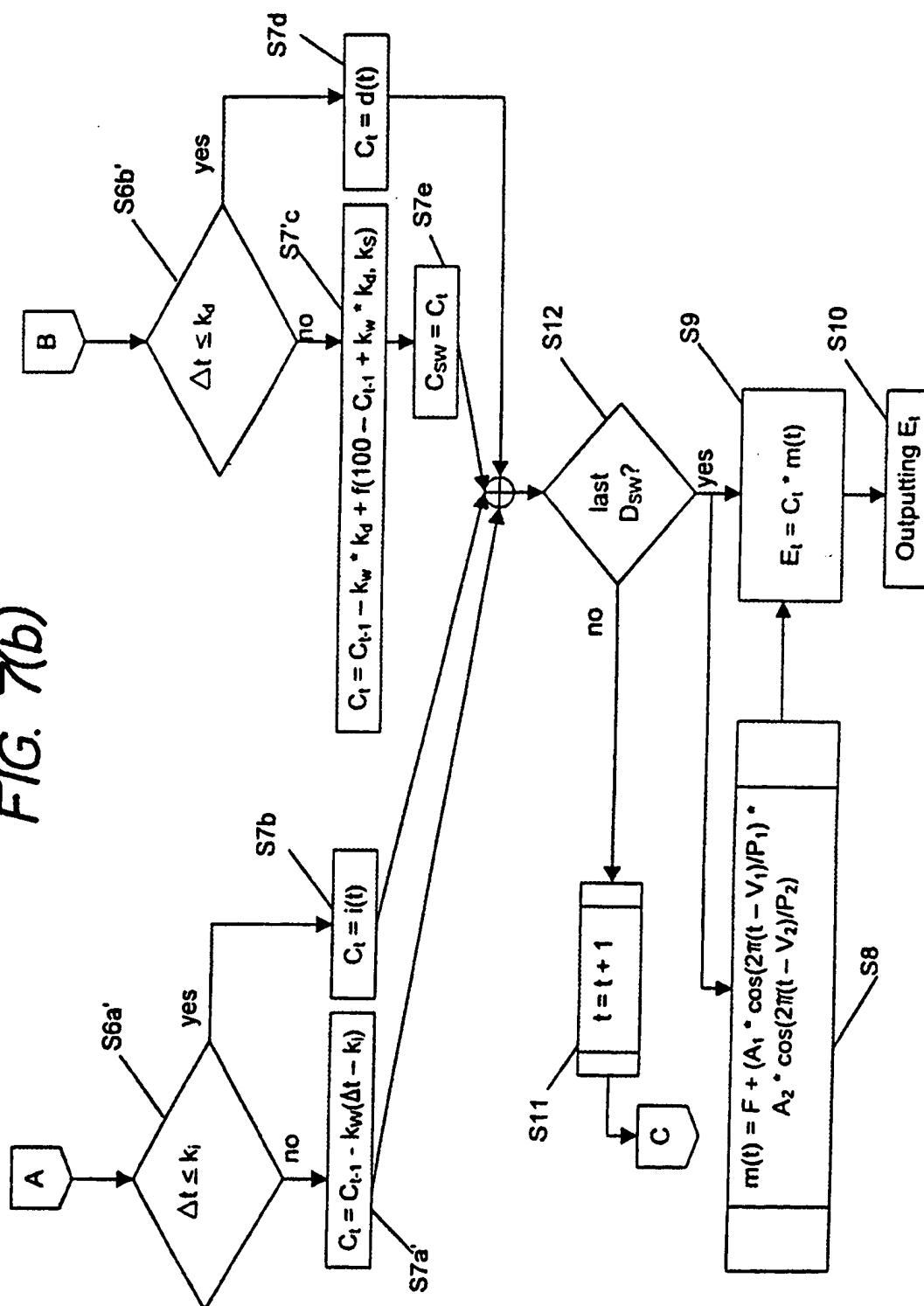
FIG. 7(a)





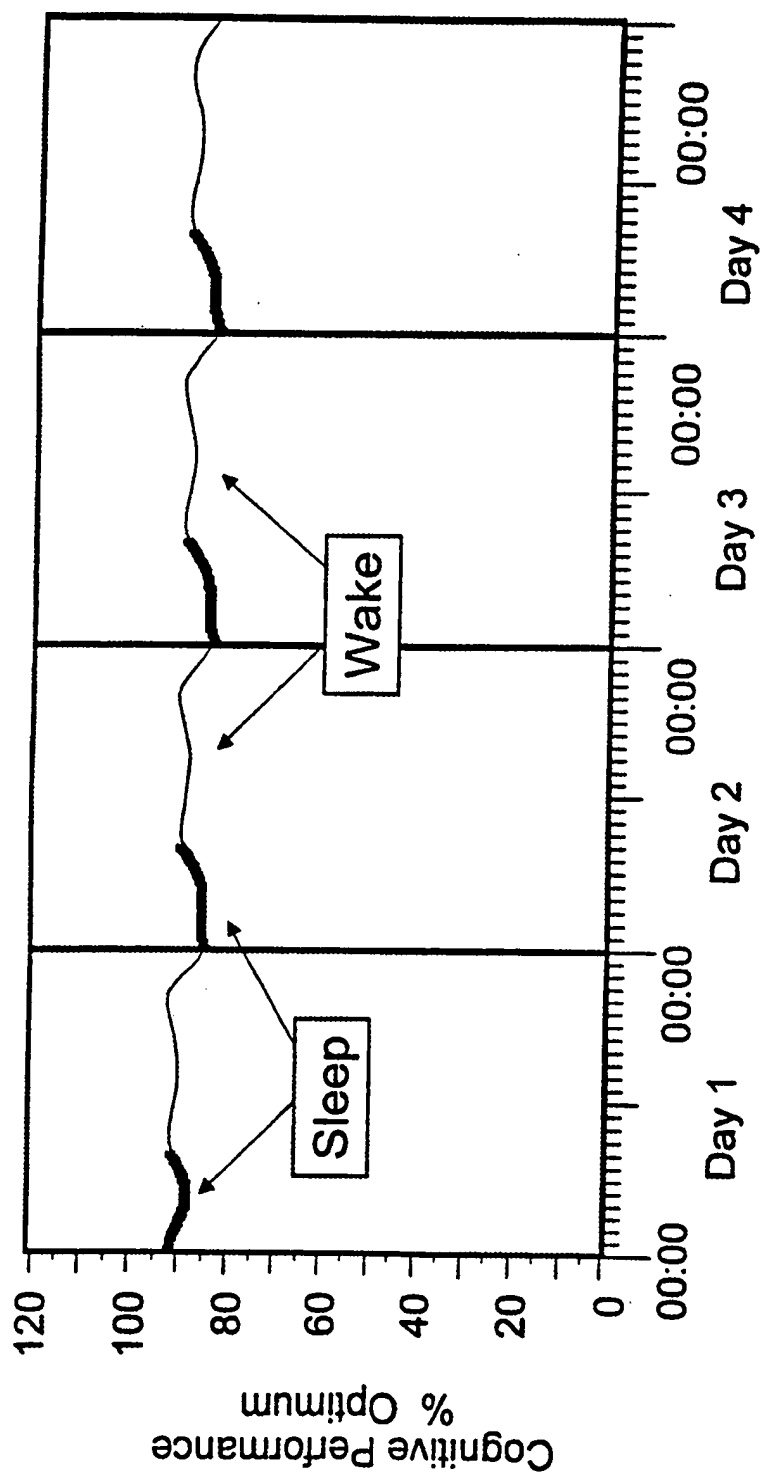
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FIG. 7(b)



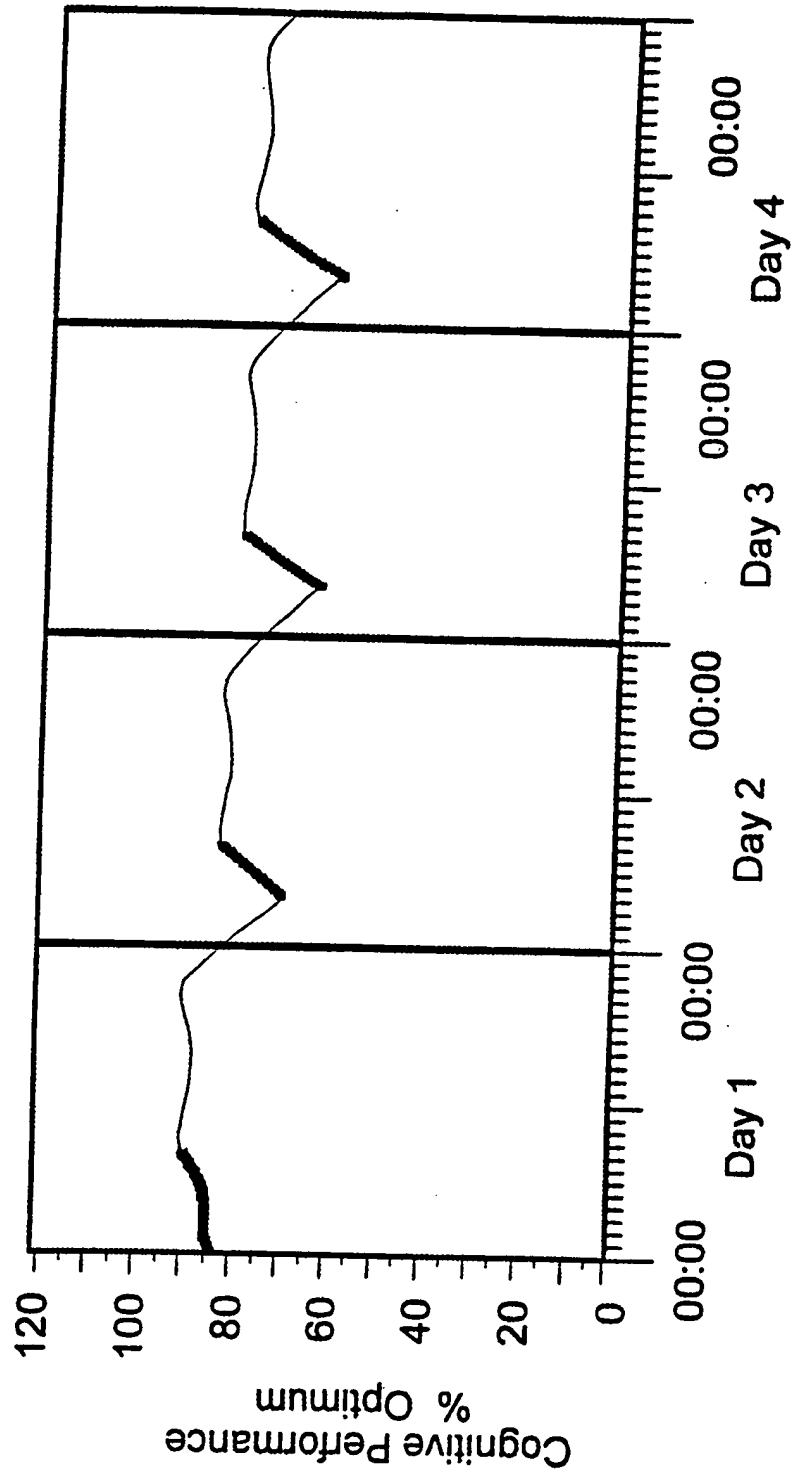
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FIG. 8



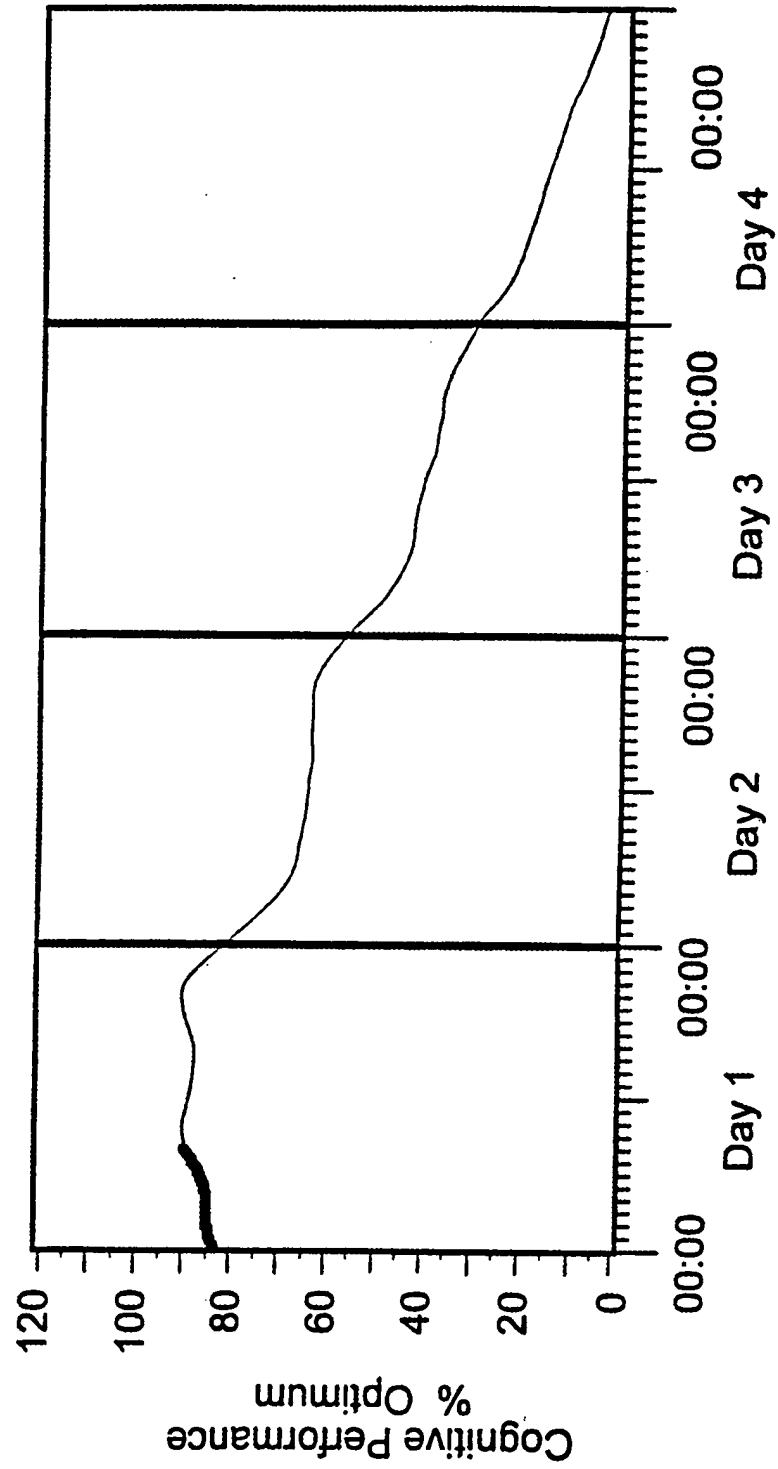
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FIG. 9



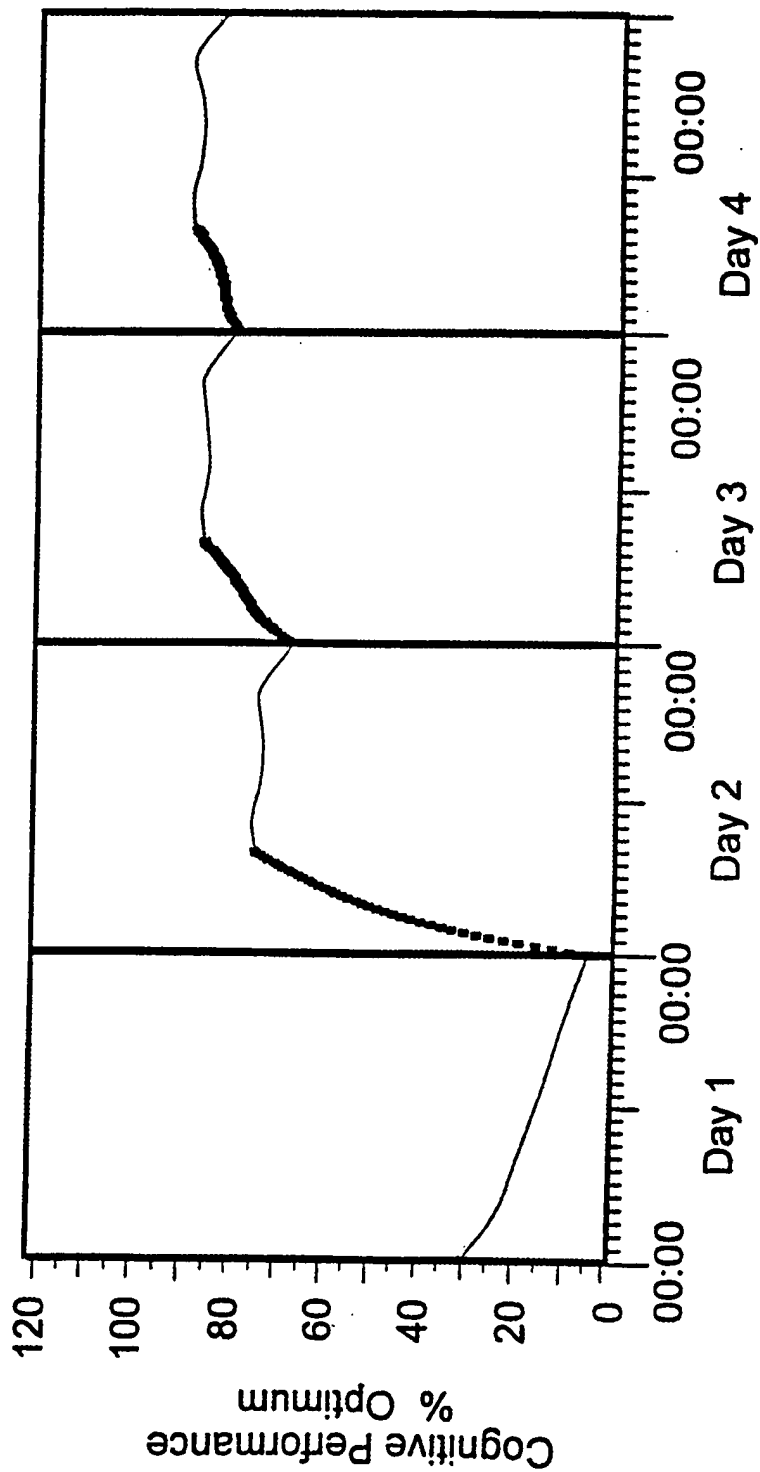
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FIG. 10



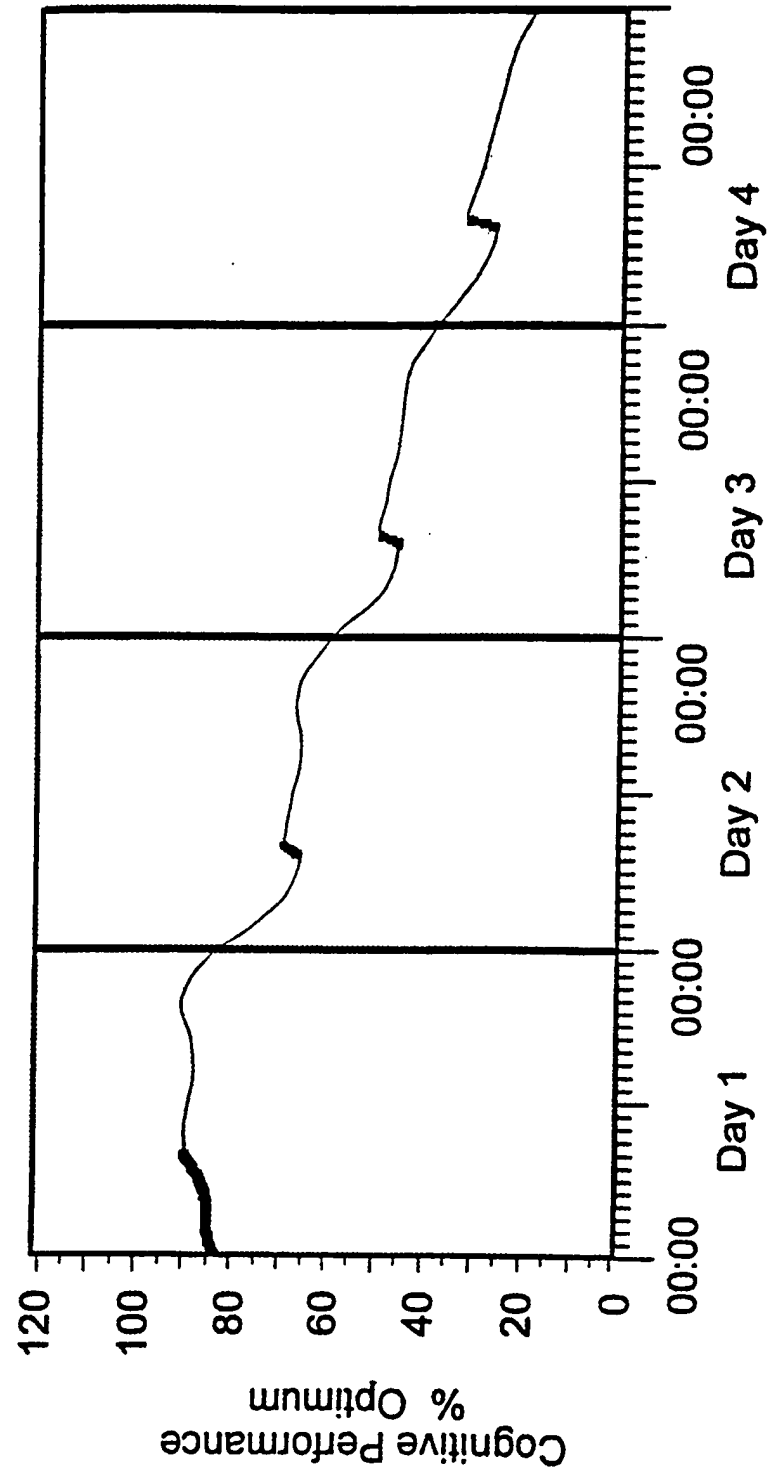
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FIG. 11



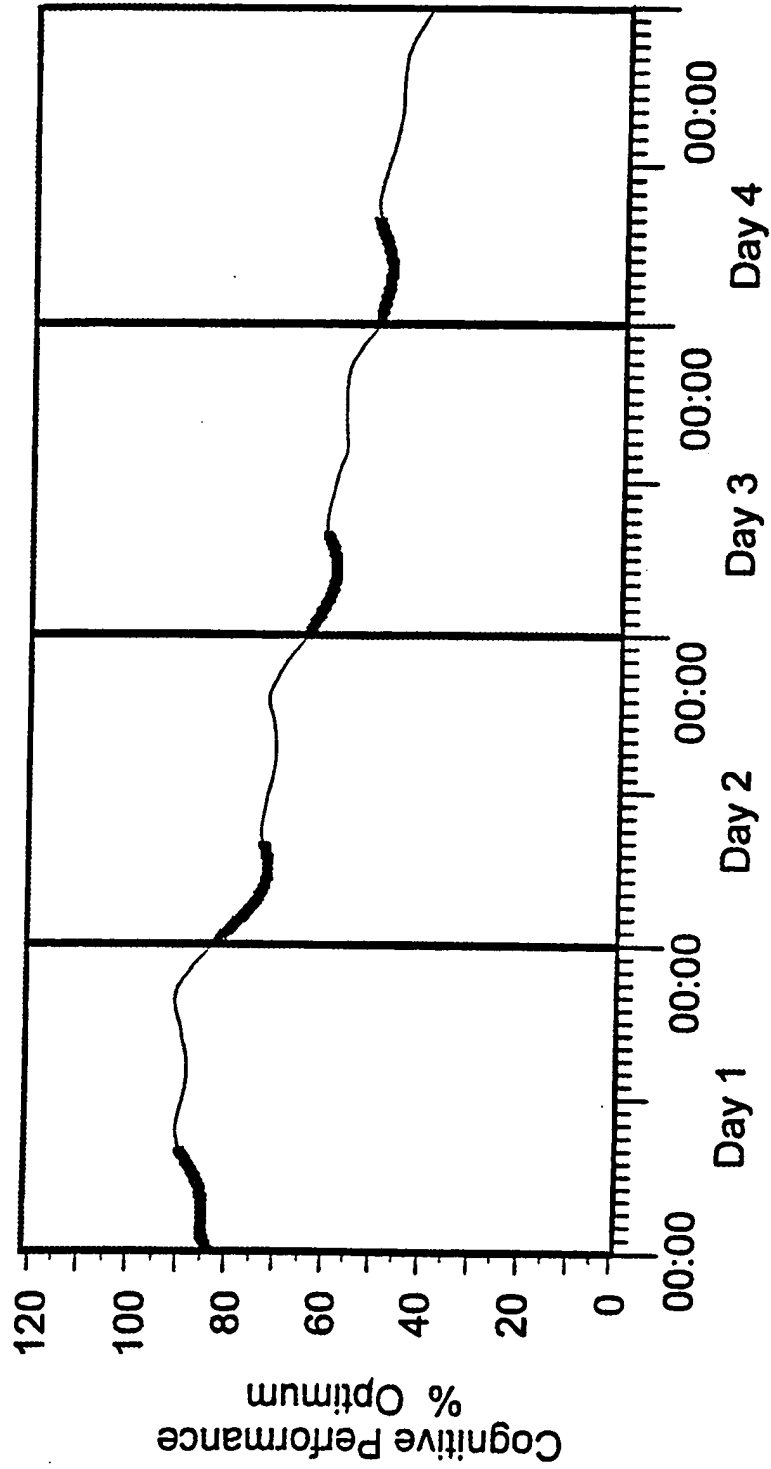
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FIG. 12



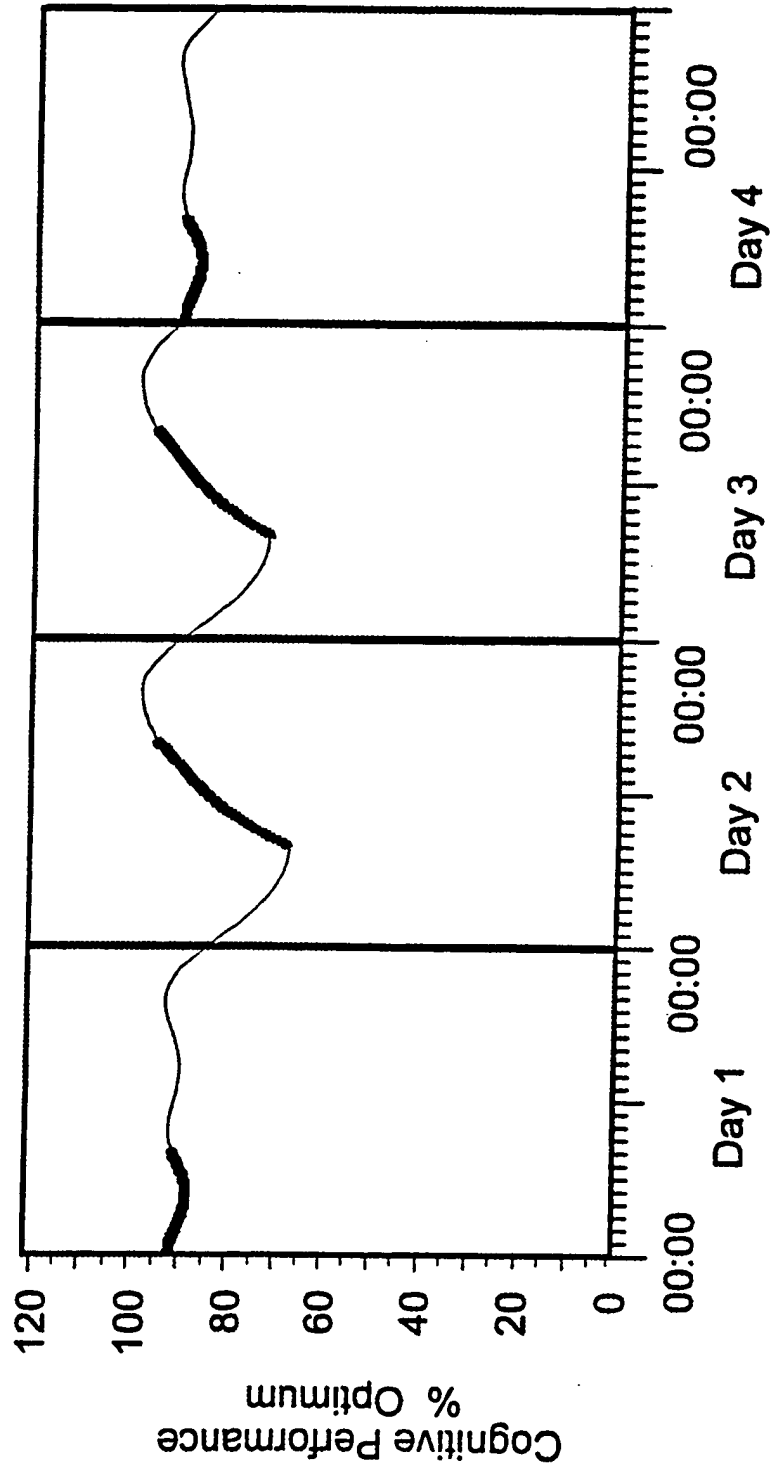
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FIG. 13



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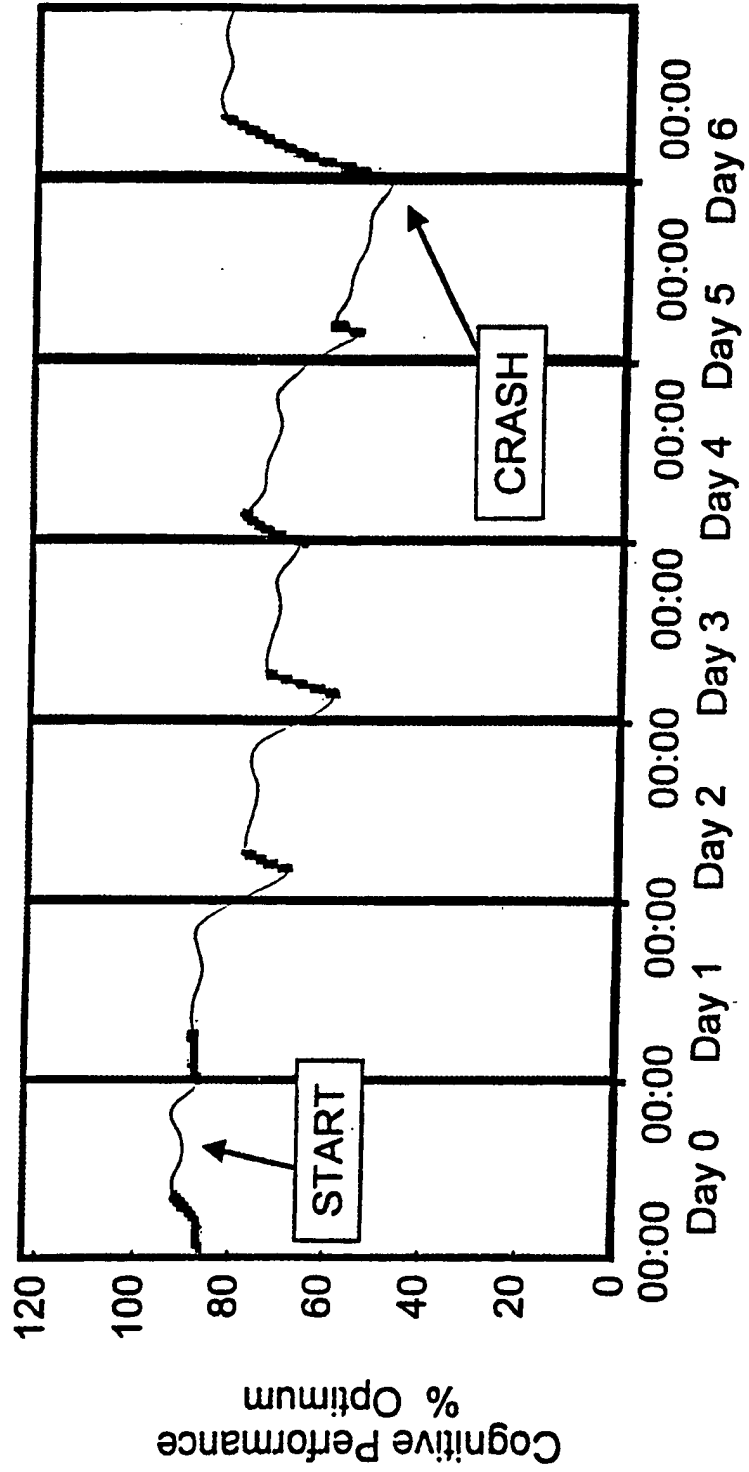
FIG. 14





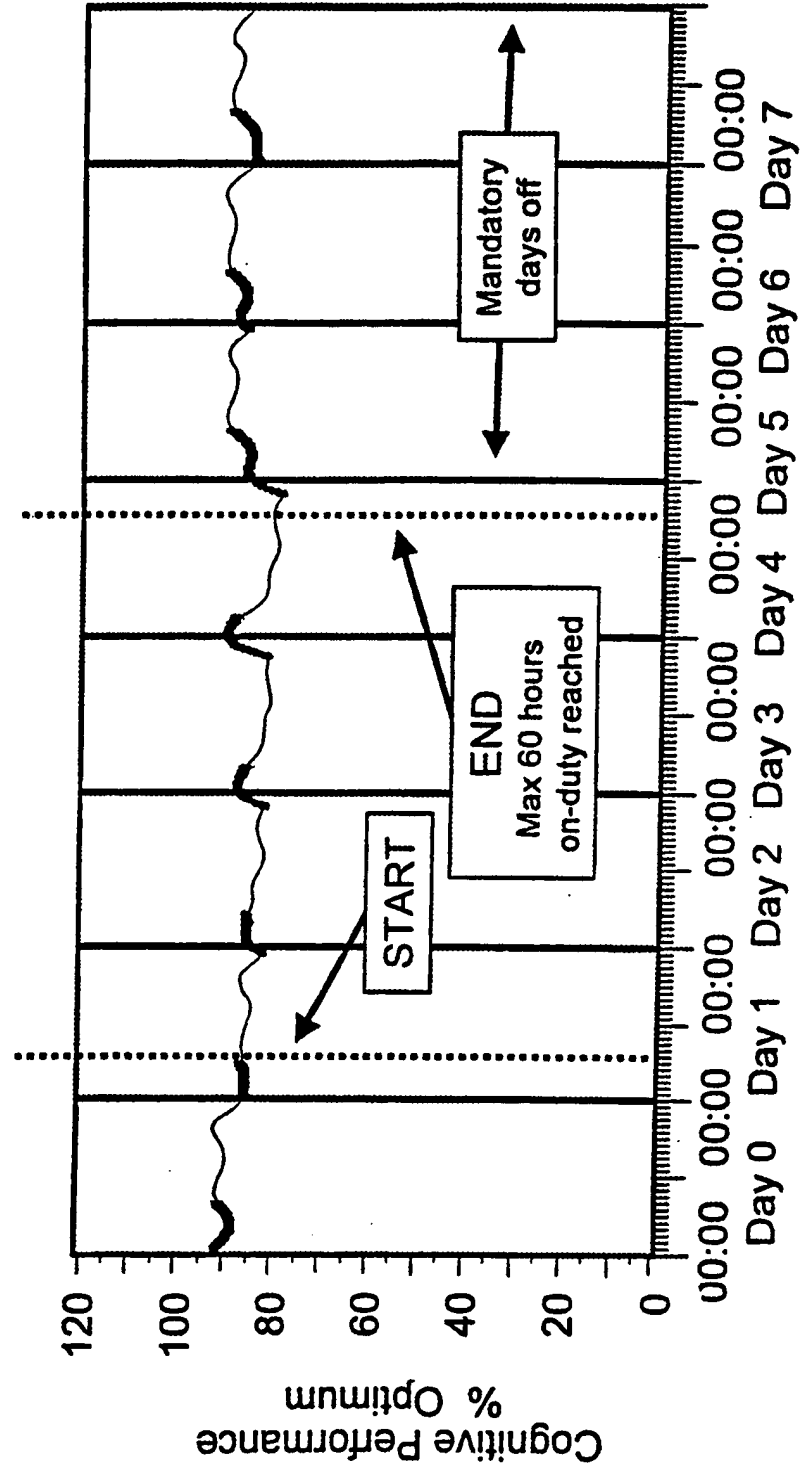
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FIG. 15



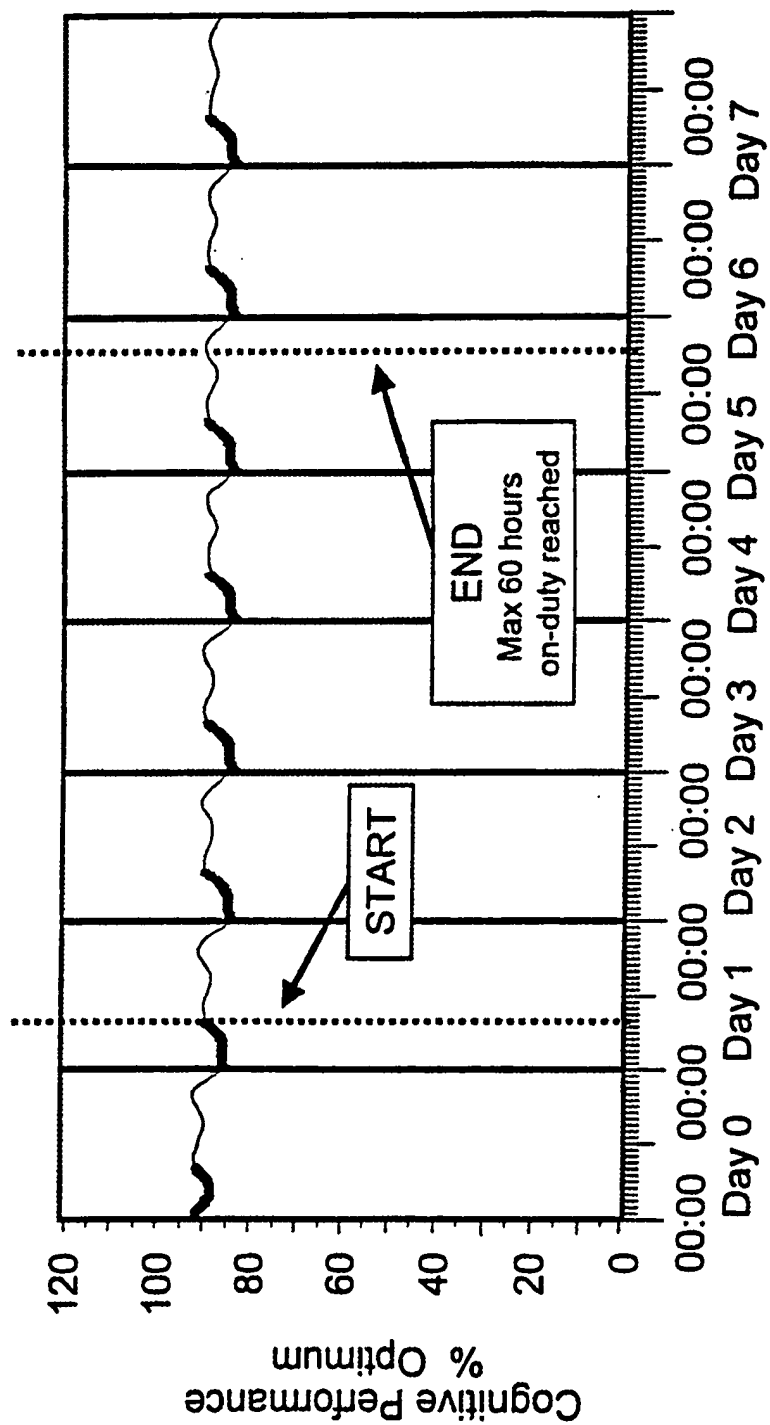
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FIG. 16

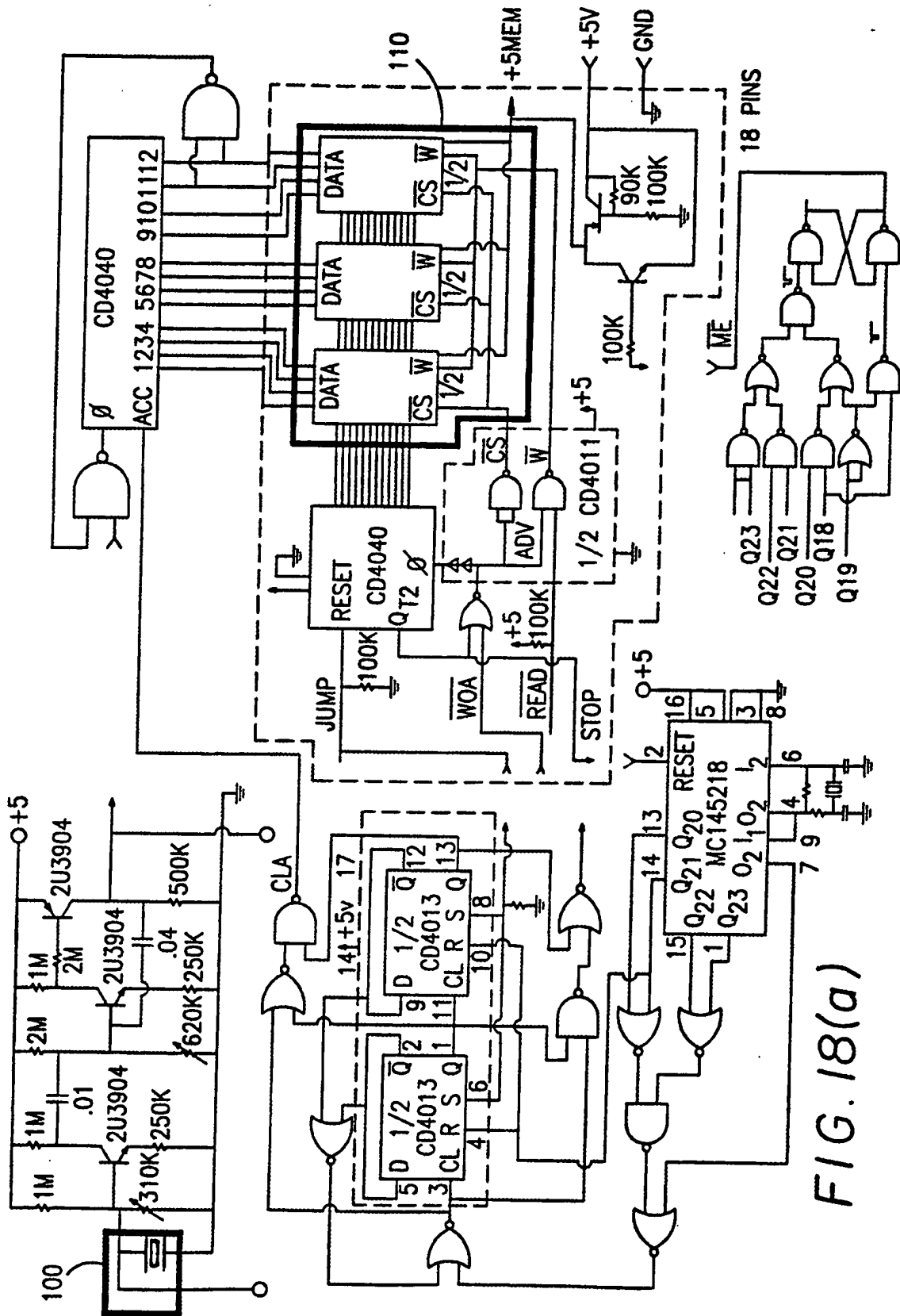


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FIG. 17



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FIG. 18(b)

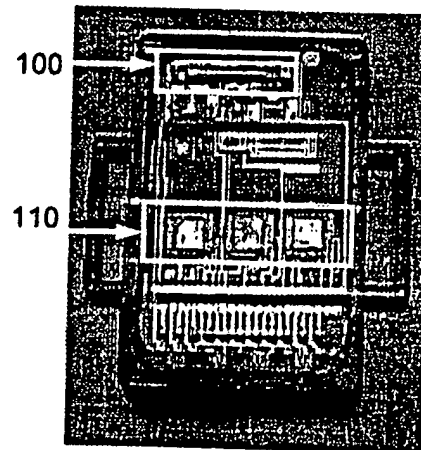


FIG. 19(b)

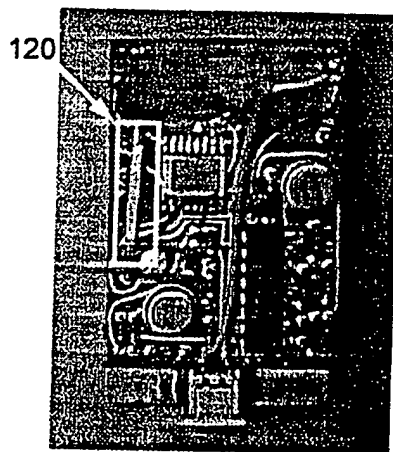
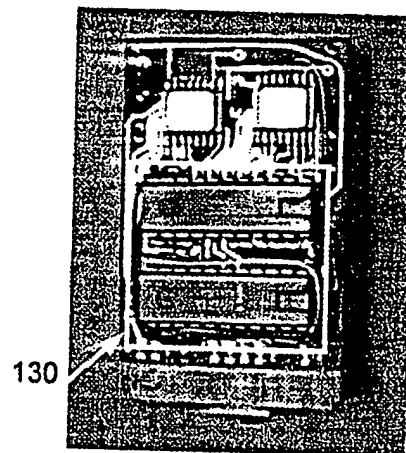
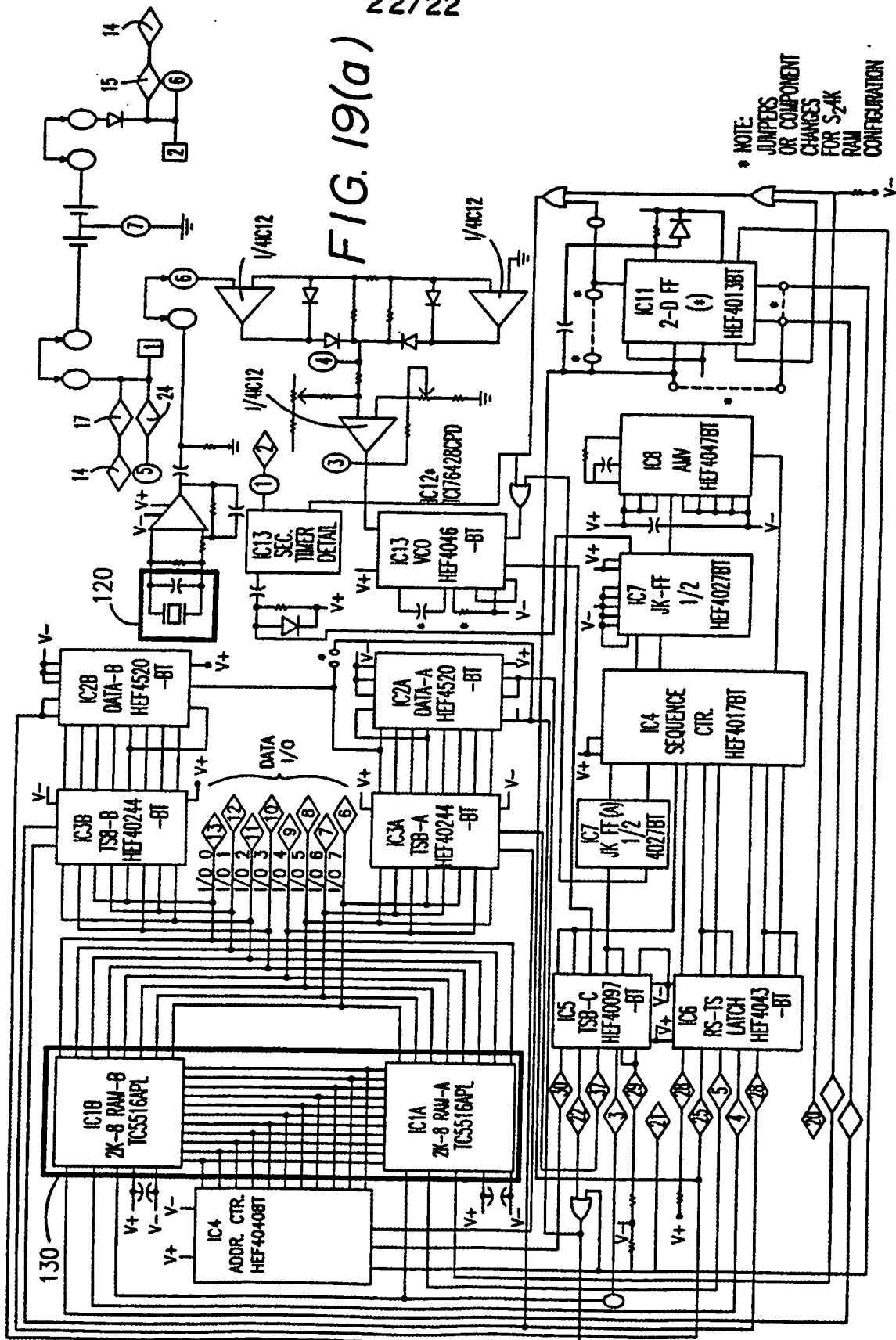


FIG. 19(c)



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FIG. 19(a)



# INTERNATIONAL SEARCH REPORT

National Application No  
**PCT/US 99/20104**

## A. CLASSIFICATION OF SUBJECT MATTER

**IPC 7 G06F19/00 G06F17/00 A61M21/00 A61B5/00**

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

**IPC 7 G06F A61M A61B**

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	<p><b>SHI Y ET AL: "Using artificial neural network for sleep/wake discrimination from wrist activity: preliminary results"</b>  <b>PROCEEDINGS OF THE 20TH ANNUAL INTERNATIONAL CONFERENCE OF THE IEEE ENGINEERING IN MEDICINE AND BIOLOGY SOCIETY. VOL.20 BIOMEDICAL ENGINEERING TOWARDS THE YEAR 2000 AND BEYOND (CAT. NO.98CH36286), HONG KONG, CHINA , 29 October 1998 (1998-10-29)</b>  <b>- 1 November 1998 (1998-11-01), pages 1113-1115 vol.3, XP002126643</b>  <b>1998, Piscataway, NJ, USA, IEEE, USA ISBN: 0-7803-5164-9</b>  <b>page 1113, left-hand column, line 1 -page 1115, left-hand column, line 2</b></p> <p style="text-align: center;">— -/-</p>	<p><b>1,14,26, 40</b></p>

☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

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- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

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- "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
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Date of the actual completion of the international search

**23 December 1999**

Date of mailing of the international search report

**11/01/2000**

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2  
NL - 2280 HV Rijswijk  
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  
Fax: (+31-70) 340-3016

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**Schenkels, P**

## INTERNATIONAL SEARCH REPORT

onal Application No

PCT/US 99/20104

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	US 5 433 223 A (MITCHELL ROSS E ET AL) 18 July 1995 (1995-07-18) cited in the application column 5, line 3 -column 6, line 40 column 9, line 1 - line 39 column 17, line 48 - line 66; figures 1-5	1,14,26, 40
A	VAN SOMEREN E J W: "Actigraphic monitoring of movement and rest-activity rhythms in aging, Alzheimer's disease, and Parkinson's disease" IEEE TRANSACTIONS ON REHABILITATION ENGINEERING, DEC. 1997, IEEE, USA, vol. 5, no. 4, pages 394-398, XP002126644 ISSN: 1063-6528 page 394, left-hand column, line 1 -page 397, right-hand column, line 29; figures 1-4	1,14,26, 40
A	KRIPKE D F ET AL: "WRIST ACTIGRAPHIC MEASURES OF SLEEP AND RHYTHMS" ELECTROENCEPHALOGRAPHY AND CLINICAL NEUROPHYSIOLOGY,NL,ELSEVIER, vol. 44, no. 5, May 1978 (1978-05), page 674-676 XP000856806 ISSN: 0013-4649 page 674, left-hand column, line 1 -page 676, left-hand column, line 25; figure 1	1,14,26, 40
A	MONK T H ET AL: "A PARALLELISM BETWEEN HUMAN BODY TEMPERATURE AND PERFORMANCE INDEPENDENT OF THE ENDOGENOUS CIRCADIAN PACEMAKER" JOURNAL OF BIOLOGICAL RHYTHMS,GB,GUILFORD PRESS, LONDON, vol. 13, no. 2, April 1998 (1998-04), page 113-122 XP000856617 ISSN: 0748-7304 page 117, right-hand column, line 34 -page 118, right-hand column, line 21	1,14,26, 40



# INTERNATIONAL SEARCH REPORT

Information on patent family members

Serial Application No

PCT/US 99/20104

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
US 5433223 A	18-07-1995	NONE	